

FORM PTO-1390 (REV. 11-2000)		U.S. DEPARTMENT OF COMMERCE PATENT AND TRADEMARK OFFICE		ATTORNEY'S DOCKET NUMBER	
<b>TRANSMITTAL LETTER TO THE UNITED STATES          DESIGNATED/ELECTED OFFICE (DO/EO/US)          CONCERNING A FILING UNDER 35 U.S.C. 371</b>				VINING PROV	
INTERNATIONAL APPLICATION NO. PCT/US99/28030		INTERNATIONAL FILING DATE 24 November 1999		U.S. APPLICATION NO. (if known, see 37 CFR 1.5) <div style="font-size: 1.5em; font-weight: bold;">09/856683</div>	
PRIORITY DATE CLAIMED 25 November 1998					
TITLE OF INVENTION VIRTUAL ENDOSCOPY WITH IMPROVED IMAGE SEGMENTATION AND LESION DETECTION					
APPLICANT(S) FOR DO/EO/US David Vining, Gordon Hunt, David Ahn, David Stelts, Yaorong Ge, Paul Hemler and Tiffany Salido					
Applicant herewith submits to the United States Designated/Elected Office (DO/EO/US) the following items and other information:					
<ol style="list-style-type: none"> <li>1. <input checked="" type="checkbox"/> This is a <b>FIRST</b> submission of items concerning a filing under 35 U.S.C. 371.</li> <li>2. <input type="checkbox"/> This is a <b>SECOND</b> or <b>SUBSEQUENT</b> submission of items concerning a filing under 35 U.S.C. 371.</li> <li>3. <input type="checkbox"/> This is an express request to begin national examination procedures (35 U.S.C. 371(f)). The submission must include items (5), (6), (9) and (21) indicated below.</li> <li>4. <input checked="" type="checkbox"/> The US has been elected by the expiration of 19 months from the priority date (Article 31).</li> <li>5. <input checked="" type="checkbox"/> A copy of the International Application as filed (35 U.S.C. 371(c)(2))           <ol style="list-style-type: none"> <li>a. <input type="checkbox"/> is attached hereto (required only if not communicated by the International Bureau).</li> <li>b. <input type="checkbox"/> has been communicated by the International Bureau.</li> <li>c. <input checked="" type="checkbox"/> is not required, as the application was filed in the United States Receiving Office (RO/US).</li> </ol> </li> <li>6. <input type="checkbox"/> An English language translation of the International Application as filed (35 U.S.C. 371(c)(2)).           <ol style="list-style-type: none"> <li>a. <input type="checkbox"/> is attached hereto.</li> <li>b. <input type="checkbox"/> has been previously submitted under 35 U.S.C. 154(d)(4).</li> </ol> </li> <li>7. <input checked="" type="checkbox"/> Amendments to the claims of the International Application under PCT Article 19 (35 U.S.C. 371(c)(3))           <ol style="list-style-type: none"> <li>a. <input type="checkbox"/> are attached hereto (required only if not communicated by the International Bureau).</li> <li>b. <input type="checkbox"/> have been communicated by the International Bureau.</li> <li>c. <input type="checkbox"/> have not been made; however, the time limit for making such amendments has NOT expired.</li> <li>d. <input checked="" type="checkbox"/> have not been made and will not be made.</li> </ol> </li> <li>8. <input type="checkbox"/> An English language translation of the amendments to the claims under PCT Article 19 (35 U.S.C. 371 (c)(3)).</li> <li>9. <input type="checkbox"/> An oath or declaration of the inventor(s) (35 U.S.C. 371(c)(4)).</li> <li>10. <input type="checkbox"/> An English language translation of the annexes of the International Preliminary Examination Report under PCT Article 36 (35 U.S.C. 371(c)(5)).</li> </ol>					
<b>Items 11 to 20 below concern document(s) or information included:</b>					
<ol style="list-style-type: none"> <li>11. <input type="checkbox"/> An Information Disclosure Statement under 37 CFR 1.97 and 1.98.</li> <li>12. <input type="checkbox"/> An assignment document for recording. A separate cover sheet in compliance with 37 CFR 3.28 and 3.31 is included.</li> <li>13. <input checked="" type="checkbox"/> A FIRST preliminary amendment.</li> <li>14. <input type="checkbox"/> A SECOND or SUBSEQUENT preliminary amendment.</li> <li>15. <input type="checkbox"/> A substitute specification.</li> <li>16. <input type="checkbox"/> A change of power of attorney and/or address letter.</li> <li>17. <input type="checkbox"/> A computer-readable form of the sequence listing in accordance with PCT Rule 13ter.2 and 35 U.S.C. 1.821 - 1.825.</li> <li>18. <input checked="" type="checkbox"/> A second copy of the published international application under 35 U.S.C. 154(d)(4).</li> <li>19. <input type="checkbox"/> A second copy of the English language translation of the international application under 35 U.S.C. 154(d)(4).</li> <li>20. <input checked="" type="checkbox"/> Other items or information: Certificate of Express Mailing (37 CFR§1.10)            Return Receipt Postcard.         </li> </ol>					

1890  
 1891  
 1892  
 1893  
 1894  
 1895  
 1896  
 1897  
 1898  
 1899  
 1900  
 1901  
 1902  
 1903  
 1904  
 1905  
 1906  
 1907  
 1908  
 1909  
 1910  
 1911  
 1912  
 1913  
 1914  
 1915  
 1916  
 1917  
 1918  
 1919  
 1920  
 1921  
 1922  
 1923  
 1924  
 1925  
 1926  
 1927  
 1928  
 1929  
 1930  
 1931  
 1932  
 1933  
 1934  
 1935  
 1936  
 1937  
 1938  
 1939  
 1940  
 1941  
 1942  
 1943  
 1944  
 1945  
 1946  
 1947  
 1948  
 1949  
 1950  
 1951  
 1952  
 1953  
 1954  
 1955  
 1956  
 1957  
 1958  
 1959  
 1960  
 1961  
 1962  
 1963  
 1964  
 1965  
 1966  
 1967  
 1968  
 1969  
 1970  
 1971  
 1972  
 1973  
 1974  
 1975  
 1976  
 1977  
 1978  
 1979  
 1980  
 1981  
 1982  
 1983  
 1984  
 1985  
 1986  
 1987  
 1988  
 1989  
 1990  
 1991  
 1992  
 1993  
 1994  
 1995  
 1996  
 1997  
 1998  
 1999  
 2000  
 2001  
 2002  
 2003  
 2004  
 2005  
 2006  
 2007  
 2008  
 2009  
 2010  
 2011  
 2012  
 2013  
 2014  
 2015  
 2016  
 2017  
 2018  
 2019  
 2020  
 2021  
 2022  
 2023  
 2024  
 2025  
 2026  
 2027  
 2028  
 2029  
 2030  
 2031  
 2032  
 2033  
 2034  
 2035  
 2036  
 2037  
 2038  
 2039  
 2040  
 2041  
 2042  
 2043  
 2044  
 2045  
 2046  
 2047  
 2048  
 2049  
 2050  
 2051  
 2052  
 2053  
 2054  
 2055  
 2056  
 2057  
 2058  
 2059  
 2060  
 2061  
 2062  
 2063  
 2064  
 2065  
 2066  
 2067  
 2068  
 2069  
 2070  
 2071  
 2072  
 2073  
 2074  
 2075  
 2076  
 2077  
 2078  
 2079  
 2080  
 2081  
 2082  
 2083  
 2084  
 2085  
 2086  
 2087  
 2088  
 2089  
 2090  
 2091  
 2092  
 2093  
 2094  
 2095  
 2096  
 2097  
 2098  
 2099  
 2100  
 2101  
 2102  
 2103  
 2104  
 2105  
 2106  
 2107  
 2108  
 2109  
 2110  
 2111  
 2112  
 2113  
 2114  
 2115  
 2116  
 2117  
 2118  
 2119  
 2120  
 2121  
 2122  
 2123  
 2124  
 2125  
 2126  
 2127  
 2128  
 2129  
 2130  
 2131  
 2132  
 2133  
 2134  
 2135  
 2136  
 2137  
 2138  
 2139  
 2140  
 2141  
 2142  
 2143  
 2144  
 2145  
 2146  
 2147  
 2148  
 2149  
 2150  
 2151  
 2152  
 2153  
 2154  
 2155  
 2156  
 2157  
 2158  
 2159  
 2160  
 2161  
 2162  
 2163  
 2164  
 2165  
 2166  
 2167  
 2168  
 2169  
 2170  
 2171  
 2172  
 2173  
 2174  
 2175  
 2176  
 2177  
 2178  
 2179  
 2180  
 2181  
 2182  
 2183  
 2184  
 2185  
 2186  
 2187  
 2188  
 2189  
 2190  
 2191  
 2192  
 2193  
 2194  
 2195  
 2196  
 2197  
 2198  
 2199  
 2200  
 2201  
 2202  
 2203  
 2204  
 2205  
 2206  
 2207  
 2208  
 2209  
 2210  
 2211  
 2212  
 2213  
 2214  
 2215  
 2216  
 2217  
 2218  
 2219  
 2220  
 2221  
 2222  
 2223  
 2224  
 2225  
 2226  
 2227  
 2228  
 2229  
 2230  
 2231  
 2232  
 2233  
 2234  
 2235  
 2236  
 2237  
 2238  
 2239  
 2240  
 2241  
 2242  
 2243  
 2244  
 2245  
 2246  
 2247  
 2248  
 2249  
 2250  
 2251  
 2252  
 2253  
 2254  
 2255  
 2256  
 2257  
 2258  
 2259  
 2260  
 2261  
 2262  
 2263  
 2264  
 2265  
 2266  
 2267  
 2268  
 2269  
 2270  
 2271  
 2272  
 2273  
 2274  
 2275  
 2276  
 2277  
 2278  
 2279  
 2280  
 2281  
 2282  
 2283  
 2284  
 2285  
 2286  
 2287  
 2288  
 2289  
 2290  
 2291  
 2292  
 2293  
 2294  
 2295  
 2296  
 2297  
 2298  
 2299  
 2300  
 2301  
 2302  
 2303  
 2304  
 2305  
 2306  
 2307  
 2308  
 2309  
 2310  
 2311  
 2312  
 2313  
 2314  
 2315  
 2316  
 2317  
 2318  
 2319  
 2320  
 2321  
 2322  
 2323  
 2324  
 2325  
 2326  
 2327  
 2328  
 2329  
 2330  
 2331  
 2332  
 2333  
 2334  
 2335  
 2336  
 2337  
 2338  
 2339  
 2340  
 2341  
 2342  
 2343  
 2344

09/856683

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re: Vining et al.

Serial No.: 09/856,683

Filed: May 23, 2001

For: VIRTUAL ENDOSCOPY WITH IMPROVED  
IMAGE SEGMENTATION AND LESION  
DETECTION

: Group Art Unit: N/A

:  
: Examiner: N/A  
:  
:  
:  
:

**PRELIMINARY AMENDMENT**

Assistant Commissioner for Patent  
Washington, D.C. 20231

Dear Sir:

Please amend the application, without prejudice, as follows:

**In the Specification**

Please change the last paragraph of page 15 which continues to page 16 to read as:

Referring again to Fig. 1, when a patient's colon is to be rendered and a contrast agent has been used to opacify stool and/or residual water, the segmentation step 35 includes the step of digitally removing the opacified stool/water from the CT image data in order to render an unimpeded colon. Digital removal of opacified feces is difficult due to the fact that simple subtraction leaves behind edge gradients. The appearance of edge gradients is illustrated in Figs. 15-17. Fig. 15 is a schematic representation of a section of image data representing hypothetical CT image data showing the measured X-ray attenuation values associated with a two dimensional array of voxels. Soft tissue is represented by a value of 100, opacified stool by a value of 200, and air by a value of 0. In this simplified model, actual interfaces between different species is presumed to lie on the boundary between adjacent voxels. Accordingly, an interface 601 between soft tissue and

09/856683-0000

opacified stool appears as the boundary between the voxels having a value of 100 and those having a value of 200. Similarly, an interface 600 between opacified stool and air appears as the boundary between the voxels having a value of 200 and those having a value of 0.

Please change the first paragraph of the Application, which was added by the first preliminary amendment, to read as:

This application claims the benefit of co-pending U.S. Provisional Application Serial No. 60/109,966, filed November 25, 1998, such application being incorporated herein by reference. This application also claims the benefit of co-pending U.S. Application Serial No. 09/299,061 filed on April 23, 1999, which is a continuation of U.S. Application Serial No. 08/805,584 filed on February 24, 1997, now issued as U.S. Patent 5,920,319.

In the Claims:

**Please add new claims:**

52. A method for guiding biopsy of a tissue by accessing the tissue via a three-dimensional structure having a lumen comprising the steps of:
  - a. providing a three-dimensional volume of data representing at least one physical property associated with the three-dimensional structure;
  - b. isolating a first region of interest of the structure having the lumen;
  - c. rendering the first region of interest in an interactive three-dimensional display;
  - d. inserting a simulated biopsy needle within the lumen of the rendered first region of interest;
  - e. determining the position of the simulated biopsy needle within the rendered first region of interest having the lumen; and
  - f. displaying the simulated biopsy needle within the rendered first region of interest having the lumen.
53. The method of claim 52 comprising the step of isolating the first region of interest from the volume of data based on a selected value of the physical property associated

with the three-dimensional structure.

54. The method of claim 53 wherein the step of isolating the first region of interest comprises segmenting the first region of interest.
55. The method of any one of claims 52 - 54 comprising:
  - providing a three-dimensional volume of data representing at least one physical property associated with the tissue;
  - isolating a second region of interest of the structure having the tissue;
  - rendering the second region of interest in the interactive three-dimensional display; and
  - creating at least one target zone of needle placement at a selected site of the rendered tissue.
56. The method of claim 55 comprising the step of isolating the second region of interest from the volume of data based on a selected value of the physical property associated with the tissue.
57. The method of claim 56 wherein the step of isolating the second region of interest comprises segmenting the second region of interest.
58. The method of claim 55 wherein the position determining step includes:
  - determining the position of the needle relative to a selected target zone;
  - generating a ready-to-strike indicator if the needle is within a desired location relative to the selected target zone; and
  - calculating a movement pathway for the needle to strike the target zone if the needle is not within a desired location relative to the selected target zone.
59. The method of claim 52 comprising:
  - providing a three-dimensional volume of data representing at least one physical property associated with mediastinal tissue;
  - isolating a second region of interest of the structure having the mediastinal tissue;
  - rendering the second region of interest in the interactive three-dimensional display; and
  - positioning the needle so as to avoid the rendered mediastinal tissue.
60. A system for interactively displaying three dimensional structures comprising:
  - a. volume formation means for forming a three-dimensional volume of data representing at least one physical property associated with a three dimensional structure;
  - b. a wireframe generator for generating a wireframe model having vertices of a region of interest representing the three-dimensional structure;
  - c. refinement means for refining the wireframe model by using a geometric deformable model; and
  - d. a renderer for rendering the wireframe model in an interactive three-dimensional display.

61. The system of claim 60 wherein the renderer produces a virtual three-dimensional environment.
62. The system of claim 60 comprising an isolator for isolating the region of interest from the volume of data based on selected values of the physical property representing the region of interest.
63. The system of claim 62 wherein the renderer produces a virtual three-dimensional environment.
64. The system of claim 62 wherein the isolator comprises a surface model identification means for identifying a surface model and wherein the wireframe generator is adapted to create a wireframe model of the surface model.
65. The system of claim 62 wherein the isolator comprises a segmentation producer for segmenting the region of interest from the volume of data.
66. The system of claim 65 wherein the isolator comprises an isosurface creation means for creating an isosurface of the isolated region of interest.
67. The system of claim 66 wherein the wireframe generator is adapted to create a wireframe model of the isosurface.
68. The system of any one of claims 60, 64, or 66 comprising:  
a vertex grouper for grouping the vertices of the wireframe model into regions having abnormal structure; and  
an analyzer for analyzing the regions having abnormal structure.
69. The system of claim 68 wherein the vertex grouper comprises:  
a mean curvature calculator for calculating a mean curvature at each of a selected number of vertices;  
a gaussian curvature calculator for calculating a gaussian curvature at each of the selected number of vertices; and  
classification means for classifying the selected vertices according to the mean curvatures and the gaussian curvatures of the vertices.
70. A computer-implemented method for interactively displaying a three-dimensional rendering of a structure having a lumen comprising the steps of:  
a. providing a three-dimensional volume of data representing at least one physical property associated with the structure having the lumen;  
b. generating a wireframe model having vertices representing a region of interest of the structure having the lumen;  
c. refining the wireframe model using a geometric deformable model; and  
d. rendering the wireframe model in an interactive three-dimensional display.
71. The method of claim 70 wherein the wireframe model comprises a plurality of vertices, the method comprising:

grouping the vertices of the wireframe model into regions having a characteristic indicating abnormal structure;  
analyzing the regions having abnormal structure to identify regions having a high degree of abnormality; and  
rendering the wireframe model in an interactive three-dimensional display to indicate the regions having abnormal structure.

72. The method of claim 71 wherein the grouping step comprises:
  - calculating a mean curvature at selected vertices;
  - calculating a gaussian curvature at the selected vertices;
  - classifying the selected vertices according to the mean curvature and the gaussian curvature; and
  - grouping adjoining selected vertices into patches of same vertex classification.
73. The method of claim 72 wherein the analyzing step comprises labeling patches having an elliptical classification as a potential lesion.
74. The method of claim 73 wherein the analyzing step comprises eliminating from the grouping selected elliptical patches which point outwardly from the lumen.
75. The method of claim 74 wherein the analyzing step comprises calculating principal curvatures of the patches from the mean and gaussian curvatures and eliminating from the grouping any elliptical patches having substantially different principal curvatures.
76. The method of claim 70 comprising calculating a normal vector for each vertex in the wireframe model.
77. The method of claim 76 wherein the refinement step comprises moving each vertex of the wireframe model along its normal vector direction at a speed that is a function of a surface curvature and a doubt level of a boundary estimate at that point.
78. The method of claim 76 wherein the refinement step uses a three-dimensional region-growing model as a starting point for the geometric deformable model.
79. The method of claim 78 wherein the refinement step uses a three-dimensional region-growing model as a starting point for the geometric deformable model.
80. The method of claim 70 wherein the step of generating a wireframe model comprises the steps of identifying a surface model of the region of interest and generating a wireframe model of the surface model.
81. The method of claim 80 comprising:
  - grouping the vertices of the wireframe model into regions having a characteristic indicating abnormal structure;
  - analyzing the regions having abnormal structure to identify regions having a high degree of abnormality; and
  - rendering the wireframe model in an interactive three-dimensional display to

indicate the regions having abnormal structure.

82. The method of claim 81 wherein the grouping step comprises:
  - calculating a mean curvature at selected vertices;
  - calculating a gaussian curvature at the selected vertices;
  - classifying the selected vertices according to the mean curvature and the gaussian curvature; and
  - grouping adjoining selected vertices into patches of same vertex classification.
83. The method of claim 82 wherein the analyzing step comprises labeling patches having an elliptical classification as a potential lesion.
84. The method of claim 83 wherein the analyzing step comprises eliminating from the grouping selected elliptical patches which point outwardly from the lumen.
85. The method of claim 84 wherein the analyzing step comprises calculating principal curvatures of the patches from the mean and gaussian curvatures and eliminating from the grouping any elliptical patches having substantially different principal curvatures.
86. The method of claim 70 comprising isolating the region of interest from the volume of data based on a selected value of the physical property representing the region of interest.
87. The method of claim 86 wherein the step of isolating comprises segmenting the region of interest.
88. The method of claim 87 comprising creating an isosurface of the region of interest.
89. The method of claim 88 wherein the step of generating a wireframe model comprises generating a wireframe model of the isosurface.
90. The method of claim 89 wherein, the wireframe model comprises a plurality of vertices, the method comprising:
  - grouping the vertices of the wireframe model into regions having a characteristic indicating abnormal structure;
  - analyzing the regions having abnormal structure to identify regions having a high degree of abnormality; and
  - rendering the wireframe model in an interactive three-dimensional display to indicate the regions having abnormal structure.
91. The method of claim 90 wherein the grouping step comprises:
  - calculating a mean curvature at selected vertices;
  - calculating a gaussian curvature at the selected vertices;
  - classifying the selected vertices according to the mean curvature and the gaussian curvature; and
  - grouping adjoining selected vertices into patches of same vertex classification.



92. The method of claim 91 wherein the analyzing step comprises labeling patches having an elliptical classification as a potential lesion.
93. The method of claim 92 wherein the analyzing step comprises eliminating from the grouping selected elliptical patches which point outwardly from the lumen.
94. The method of claim 93 wherein the analyzing step comprises calculating principal curvatures of the patches from the mean and gaussian curvatures and eliminating from the grouping any elliptical patches having substantially different principal curvatures.
95. A computer-implemented method for interactively displaying a three-dimensional rendering of a structure having a lumen comprising the steps of:
- providing a three-dimensional volume of data representing at least one physical property associated with the structure having the lumen;
  - generating a wireframe model having a plurality of vertices to represent the structure having the lumen;
  - grouping the vertices by:
    - calculating a mean curvature at each of a selected number of vertices,
    - calculating a gaussian curvature at each of the selected number of vertices,
    - classifying the selected vertices according to the mean curvatures and the gaussian curvatures of the vertices, and
    - collecting adjoining vertices of similar classification into patches of similar vertex classification; and
  - rendering the wireframe model in an interactive three-dimensional display.
96. The method of claim 95 comprising isolating a region of interest from the volume of data based on a selected value of the physical property representing the region of interest.
97. The method of claim 96 wherein the step of isolating comprises segmenting the region of interest.
98. The method of claim 95 wherein the step of generating a wireframe model comprises the steps of identifying a surface model of the region of interest and generating a wireframe model of the surface model.
99. The method of claim 95 comprising creating an isosurface of the region of interest.
100. The method of claim 95 comprising labeling patches having an elliptical classification as a potential lesion.
101. The method of claim 100 comprising eliminating from the grouping selected elliptical patches which point outwardly from the lumen.
102. The method of claim 101 comprising calculating principal curvatures of the patches from the mean and gaussian curvatures and eliminating from the grouping any

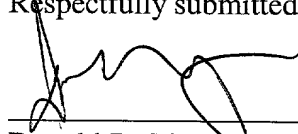
elliptical patches having substantially different principal curvatures.

103. A system for interactively displaying three dimensional structures comprising:
  - a. volume formation means for forming a three-dimensional volume of data representing at least one physical property associated with a three dimensional structure;
  - b. a wireframe generator for generating a wireframe model having a plurality of vertices to represent a region of interest of the three-dimensional structure;
  - c. a vertex grouper for collecting adjoining vertices having similar classifications based on mean and gaussian curvatures into patches of similar vertex classification; and
  - d. a renderer for rendering the wireframe model in an interactive three-dimensional display.
104. The system of claim 103 wherein the renderer produces a virtual three-dimensional environment.
105. The system of claim 103 comprising an isolator for isolating the region of interest from the volume of data based on selected values of the physical property representing the region of interest.
106. The system of claim 105 wherein the renderer produces a virtual three-dimensional environment.
107. The system of claims 105 or 106 wherein the isolator comprises a segmentation producer for segmenting the region of interest from the volume of data.
108. The system of claim 105 wherein the isolator comprises a surface model identification means for identifying a surface model and wherein the wireframe generator is adapted to create a wireframe model of the surface model.
109. The system of claim 105 wherein the isolator comprises an isosurface creation means for creating an isosurface of the isolated region of interest.
110. The system of claim 103 wherein the vertex grouper comprises:
  - a mean curvature calculator for calculating a mean curvature at each of a selected number of vertices;
  - a gaussian curvature calculator for calculating a gaussian curvature at each of the selected number of vertices; and
  - classification means for classifying the selected vertices according to the mean curvatures and the gaussian curvatures of the vertices.

**REMARKS**

Applicants have added new claims 52-110 to claim additional features of Applicants' invention. Applicants have also amended the specification at page 15 to make the recitation of reference numerals 600 and 601 consistent with the usage of such reference numerals in Figure 15. No new matter has been added. Applicants have added a claim to additional related applications.

Respectfully submitted,



Donald R. Piper, Jr.  
PTO Reg. No. 29,337

DANN DORFMAN HERRELL & SKILLMAN, PC  
1601 Market Street, Suite 720  
Philadelphia, PA 19103  
Phone: (215) 563-4100  
Fax: (215) 563-4044

0985653 0324  
10220 0395860

## ATTACHMENT

The following is a marked up version of the first paragraph of the Application:

This application claims the benefit of U.S. Provisional Application Serial No. 60/109,966, filed November 25, 1998, such application being incorporated herein by reference. This application also claims the benefit of co-pending U.S. Application Serial No. 09/299,061 filed on April 23, 1999, which is a continuation of U.S. Application Serial No. 08/805,584 filed on February 24, 1997, now issued as U.S. Patent 5,920,319.

The following is a marked up version of the last paragraph of page 15 which continues to page 16:

Referring again to Fig. 1, when a patient's colon is to be rendered and a contrast agent has been used to opacify stool and/or residual water, the segmentation step 35 includes the step of digitally removing the opacified stool/water from the CT image data in order to render an unimpeded colon. Digital removal of opacified feces is difficult due to the fact that simple subtraction leaves behind edge gradients. The appearance of edge gradients is illustrated in Figs. 15-17. Fig. 15 is a schematic representation of a section of image data representing hypothetical CT image data showing the measured X-ray attenuation values associated with a two dimensional array of voxels. Soft tissue is represented by a value of 100, opacified stool by a value of 200, and air by a value of 0. In this simplified model, actual interfaces between different species is presumed to lie on the boundary between adjacent voxels. Accordingly, an interface [600] 601 between soft tissue and opacified stool appears as the boundary between the voxels having a value of 100 and those having a value of 200. Similarly, an interface [601] 600 between opacified stool and air appears as the boundary between the voxels having a value of 200 and those having a value of 0.

# DANN, DORFMAN, HERRELL AND SKILLMAN,

A PROFESSIONAL CORPORATION

09/856683

1601 MARKET STREET • SUITE 720 • PHILADELPHIA, PA • 19102-2207

PHONE (215) 563-4100 • FAX (215) 563-4044

JC18 Rec'd PCT/PTO 23 MAY 2001

COMMISSIONER OF PATENTS AND TRADEMARKS WASHINGTON, DC 20231	DATE OF MAILING  May 23, 2001
BOX PCT	APPLICANT'S OR AGENT'S FILE REF.  VININGPROV
IDENTIFICATION OF THE INTERNATIONAL APPLICATION	
INTERNATIONAL APPLICATION NO.  PCT/US99/28030	INTERNATIONAL FILING DATE  24 NOV 99

## PRELIMINARY AMENDMENT

Please amend the specification of the above identified application follows:

Before the Field of the Invention, insert the following:

## RELATED APPLICATIONS

This application claims the benefit of U.S. Provisional Application Serial No. 60/109,966, filed November 25, 1998, such application being incorporated herein by reference.--

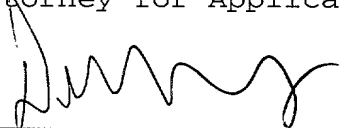
## REMARKS

The application has been amended to contain a specific reference to an earlier filed application. No new matter has been added.

Respectfully submitted,

DANN, DORFMAN, HERRELL AND SKILLMAN  
A Professional Corporation  
Attorney for Applicant

By:

  
DONALD R. PIPER, JR.  
PTO Registration No. 29,337

Telephone: 215/563-4100  
Facsimile: 215/563-4044

Rec'd PCT/PTO 23 AUG 2001 #4  
09/856683

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re the Application of :  
VINING et al. : Examiner:  
Application No. 09/856,683 : Group Art Unit:  
Filed: May 23, 2001 :  
For: Virtual Endoscopy with :  
Improved Image Segmentation :  
and Lesion Detection :

THIRD PRELIMINARY AMENDMENT

Please change the Related Applications section of the above-identified application to read as follows:

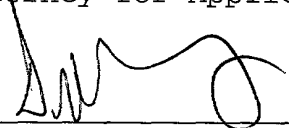
This application claims the benefit of U.S. Provisional Application Serial No. 60/109,966, filed November 25, 1998, such application being incorporated herein by reference. This application also claims the benefit of and is a continuation-in-part of co-pending U.S. Application Serial No. 09/299,061 filed on April 23, 1999, which is a continuation of U.S. Application Serial No. 08/805,584 filed on February 24, 1997, now issued as U.S. Patent 5,920,319.

Remarks

The Related Application section of the specification has been amended contain editorial revisions. No new matter has been added.

Respectfully submitted,

DANN, DORFMAN, HERRELL AND SKILLMAN  
A Professional Corporation  
Attorney for Applicant(s)

By   
DONALD R. PIPER, JR., ESQ.  
PTO Registration No. 29,337

Telephone: 215/563-4100  
Facsimile: 215/563-4044

**SECRET**

**SECRET**

**SECRET**

12/PRTS

09/856683

JC18 Rec'd PCT/PTO 23 MAY 2001

WO 00/32106

PCT/US99/28030

Virtual Endoscopy with Improved Image Segmentation  
and Lesion Detection

David J. Vining  
Gordon W. Hunt  
David K. Ahn  
David R. Stelts  
Yaorong Ge  
Paul F. Hemler  
Tiffany W. Salido

FIELD OF THE INVENTION

The present invention relates to a system and method for interactively displaying a three-dimensional rendering of a structure and, more particularly, to a system and method for automatically analyzing such structures for potential abnormalities.

BACKGROUND OF THE INVENTION

For many forms of cancer, early detection is essential for a favorable prognosis. A cancerous growth must be detected at an early stage before the cancer is allowed to grow and spread. This is particularly true for colorectal and lung cancers. As a result, endoscopic techniques have been developed to examine the colon and tracheobronchial airways for the growth of precancerous and cancerous masses.

Regarding colorectal cancer, the American Cancer Society and the National Cancer Institute recommend routine screening in order to provide a means for early detection of abnormal growths. Under those guidelines over 50 million Americans should undergo annual colorectal screening. However, only a small percentage of the population undergoes colonoscopy



screening each year. The underutilization of such screening exists, at least in part, because conventional colonoscopies require a patient to be sedated so that a flexible endoscope can be inserted into the patient's anus. Another problem is that conventional colonoscopies fail to provide access to the entire colon in approximately 15% of cases. Colonoscopies also expose patients to the risk of bowel perforation. Accordingly, it is understandable that patients are reluctant to undergo, or repeatedly subject themselves to, such an invasive procedure.

Consequently, virtual endoscopic techniques have been, and are continuing to be developed. Virtual endoscopy that utilizes computer reformation of radiologic cross-sectional images is a minimally invasive alternative to conventional fiberoptic endoscopy. Virtual endoscopy reduces the risk of perforation, does not require any sedation, and is considerably less expensive than the fiberoptic endoscopy method. For example, virtual colonoscopy techniques generally require bowel cleansing, gas distension of the colon, a 10-60 second helical computed tomography (CT) scan of a patient's abdomen and pelvis, and human visual analysis of multi-planar two-dimensional (2D) and three-dimensional (3D) images created from CT data.

Although virtual colonoscopy techniques provide excellent images of the colon in three-dimensions, a correct diagnosis relies upon a physician's ability to properly identify small (approximately 3-10 mm), and sometimes subtle, masses within hundreds of multiplanar two-dimensional and three-dimensional images. Such human inspection is time consuming,

tedious, expensive, and under certain circumstances prone to error of interpretation. Accordingly, it would be highly beneficial to provide an automatic virtual endoscopic system and method for automatically analyzing and/or detecting abnormalities, such as abnormal growths, in a targeted structure or organ system, such as the walls of a colon or trachea.

#### SUMMARY OF THE INVENTION

In accordance with the present invention, a computer-implemented method and computer system are provided for interactively displaying a three-dimensional rendering of a structure, generally having a lumen. In a specific application, the method may be utilized for analyzing regions having certain characteristics of wall structure, such as thickness or shape, to detect, for example, abnormal masses. In accordance with the method of the present invention, a three-dimensional volume of data is formed from a series of two-dimensional images representing at least one physical property associated with the three-dimensional object. A surface of a selected region of interest, optionally an isosurface, is created in the computer from the volume of data based on selected values of the physical properties representing the selected region of interest. A wireframe model of the surface is generated by the computer. The wireframe model may be purposefully deformed by the computer to more accurately match the object of interest. The wireframe model is analyzed to detect, by computer, those sections of the object having the selected characteristic such as abnormal wall structure. For example, the wireframe model includes a plurality of

vertices. The computer-implemented method groups the vertices of the wireframe model into populations having a characteristic indicating abnormal wall structure. The wireframe model with highlighted abnormal portions is then rendered by the computer into an interactive three-dimensional display on the computer monitor. Alternatively, the locations of selected abnormal portions may be stored and indexed in the computer without necessity of an interactive display.

In specific applications, a system for interactively displaying three-dimensional structures is provided. Volume formation means is included for forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with a three-dimensional body. A wireframe modeling means is provided for producing a wireframe model of a selected region or object of interest. A wireframe deforming means is provided for deforming the wireframe model to more accurately represent the object of interest. Finally, rendering means may optionally be provided for rendering the wireframe model in an interactive three-dimensional display, therein producing a virtual three-dimensional environment.

In a specific embodiment, segmentation means may be provided for segmenting the object of interest from the volume of data based on selected values of the physical property representing the region of interest. In addition, the modeling means may function to produce the wireframe model from the segmented region of interest.

5 The wireframe model may be produced to include a plurality of vertices, with each vertex having a coordinate position. In such an embodiment, the deforming means may include refinement means for refining the wireframe model by adjusting the coordinate positions of the vertices to more accurately represent the object of interest.

10 As a further option, the vertices of the wireframe model may be grouped into regions having a characteristic indicating abnormal structure. The regions of abnormal structure may then be analyzed to identify regions having a high degree of abnormality. The abnormal structure can be designated and indexed by the computer either as a comprehensive listing or  
15 as a listing dependent on the degree of abnormality.

20 The system may also include provision for targeting abnormalities and for determining the position of a biopsy needle within the three-dimensional virtual environment. In such an embodiment, an interactive three-dimensional display of a rendered wireframe model of a selected object of interest is displayed from the perspective indicating the position of the biopsy needle within the three-dimensional virtual environment.

#### 25 BRIEF DESCRIPTION OF THE DRAWINGS

30 The foregoing summary, as well as the following detailed description of the preferred embodiments of the present invention, will be better understood when read in conjunction with the accompanying drawings, in which:

Fig. 1 is a flowchart representing a method in accordance with the present invention of producing an

interactive, three-dimensional rendering of a selected organ system, such as an organ containing a lumen, and of detecting potentially abnormal areas of the organ system;

5           Fig. 2 is a flowchart representing a method for analyzing the shape of a population representing a potentially abnormal area within the selected organ system;

10           Figs. 3 a-b are graphical representations of a method for treating evolving surfaces as the level surfaces of an evolving scalar function;

          Fig. 4 is a block diagram of a system used in the method of the present invention;

15           Fig. 5 is a schematic representation of a two-dimensional plane through a wireframe model of an area of abnormal wall structure illustrating a method for determining the convexity of the area;

20           Figs. 6 a-d are diagrams illustrating methods of calculating wall thickness at a vertex of a wireframe model by analyzing the intensity profile along the normal vector originated at the vertex;

25           Fig. 7 is a schematic view of a display window for displaying a volume rendered image, a surface rendered image, and three multiplanar images along three different planes of a selected organ system;

          Fig. 8 is a schematic diagram depicting the cross-sectional area of a lumen defined by a plane through the lumen;

30           Figs. 9 a-d are diagrams representing methodology for calculating the cross sectional area of the lumen defined by the cutting plane shown in Fig. 8;

          Fig. 10 is a schematic view of a display window for displaying a volume rendered image, a surface

rendered image, and three multiplanar images along three different planes of a selected organ system combined in a single window display;

Fig. 11 is a block diagram of an alternate embodiment of the system used in the method of the present invention;

Fig. 12 is a flowchart representing a method for creating a selected surface of a region of interest;

Fig. 13 is a flowchart representing an alternate method for analyzing the shape of a population;

Fig. 14 is a flowchart representing a method for guiding endoscopic biopsy procedures in accordance with the present invention;

Fig. 15 is a schematic representation of section of two-dimensional image data containing hypothetical soft tissue/opacified stool and opacified stool/air interfaces;

Fig. 16 is a schematic representation of a simplified section of two-dimensional image data containing soft tissue/opacified stool and opacified stool/air interfaces; and

Fig. 17 is a schematic representation of the section of two-dimensional image data of Fig. 16 showing edge gradient effects resulting from a simple digital subtraction scheme.

#### DETAILED DESCRIPTION OF THE PREFERRED EMBODIMENTS

The present invention generally relates to a computer-implemented method and system, as schematically represented in Figs. 1, 2, and 4 for generating interactive, three-dimensional renderings

of three-dimensional structures generally having a lumen. The structures are usually in the general form of selected regions of a body and in particular, human or animal body organs which have hollow lumens such as colons, blood vessels, and airways. In accordance with the present invention, the interactive three-dimensional renderings are generated in a computer-controlled process from a series of two-dimensional, cross-sectional images of the selected body organ acquired, for example, from a helical computed tomography (CT) scan. The three-dimensional renderings are interactive in that such renderings can be manipulated by a user on a visual display of a computer system, such as a computer monitor, to enable movement in, around and through the three-dimensional structure, or slices or cuts of such three-dimensional structure, while simultaneously displaying multiplanar views centered on a point that changes in response to the movement, or on a point selected by the user.

The computer-generated interactive three-dimensional renderings can be constructed from surface renderings of the organ system or from volume renderings of the organ system and surrounding anatomy and may be presented from surface and volume renderings simultaneously in the same three-dimensional display window. In a selected embodiment, a surface rendering of an organ can be incorporated into a volume rendering of the surrounding anatomy using texture memory in order to achieve a combined surface/volume rendering, or a hybrid rendering effect, wherein the surface rendered organ may be highlighted in the context of the volume-rendered surrounding anatomy. In yet another embodiment,

multiplanar views such as orthogonal (sagittal, coronal, and axial) or oblique views, may be presented simultaneously with a surface-rendered organ and surrounding volume-rendered anatomy in either separate  
5 three-dimensional display windows or in a combined single three-dimensional display or window.

A computer-controlled system 20 in accordance with the present invention is shown schematically in Fig. 4. The system 20 is generally as described in  
10 U.S. Patent No. 5,782,762, which is incorporated herein by reference. Briefly, a computer console 24 is used to operate a scanner 22 to produce a series of two-dimensional, cross-sectional images of a selected three-dimensional object. The two-dimensional images  
15 are transferred from the console 24 to a computer graphics workstation 26 over a computer network 25. The graphics computer 26 is used to generate a three-dimensional rendering, such as a three dimensional surface rendering, of the selected object and to automatically identify potentially abnormal regions of the structure. The three-dimensional rendering is  
20 displayed on a computer monitor 28. Additionally, the displayed rendering can be recorded on a video recorder 30 or photographed and/or digitally recorded for future viewing. Various inputs, such as a  
25 computer mouse 27, are provided to the graphics computer to permit a user to manipulate the displayed imagery.

The scanner 22 can be a General Electric HiSpeed Advantage Helical CT Scanner (GE Medical Systems, Milwaukee, WI). For scanning a patient's lungs, the scanner is typically operated with an X-ray beam collimation of 3 mm, a pitch of 1:1, a 1 mm  
30



reconstruction interval, a display field-of-view of 25.6 cm, and a 512x512 matrix. For scanning a patient's colon, the scanner is typically operated with a beam collimation of 5 mm, a pitch of 2:1, a 1 mm reconstruction interval, a display field-of-view of 40 cm, and a 512x512 matrix. These protocols typically result in approximately 200 thoracic CT images or 500 abdominal CT images, which occupy approximately 100 MB and 250 MB of disk storage space, respectively. However, the protocols can be modified to optimally acquire image data of specific organ systems, or to adapt to advances in scanner technology. Alternatively, MR scanning can be employed, or any future technology that generates image data.

The image data can be stored in any of a variety of image formats. For example, the image data can be stored using the digital imaging and communications in medicine (DICOM) standard, or as raw binary slices, or in a variety of volume formats. For example, the image data can be stored in the computer memory in an internal data format which allows the image files to be saved as a single data volume instead of individual image files. The internal data format can also permit standard compression and uncompression techniques to be incorporated, thereby reducing computer disk storage requirements.

An alternate arrangement of the system in accordance with the present invention is shown schematically in Fig. 11. The system of Fig. 11 is particularly well suited for use with a centralized high-speed graphics computer. The scanner 422, computer console 424, graphics computer 426, memory

412, mouse 427, monitor 428, and video recorder 430 are essentially the same as the corresponding components of system 20. However, in system 420, the mouse 427, monitor 428, and video recorder 430 are connected to computer console 424. Additionally, the graphics computer 426 is connected to the computer console 424 via a computer network 429 such as an outside network permitting remote connection of the graphics computer. The computer network 429 enables the graphics computer 426 to both receive image data from the computer console 424 and transmit the results of subsequent image data analysis to the computer console 424. Accordingly, the graphics computer 426 can be located remotely from computer console 424. Further, graphics computer 426 can be used to analyze image data received from more than one computer console.

A method for generating interactive, three-dimensional renderings of a selected structure having a lumen and indicating potentially abnormal regions of the structure in accordance with the present invention is generally set forth in Fig. 1. The method can be implemented by computer. The steps of patient preparation 32, acquisition of two-dimensional images 33, formation of a three-dimensional volume 34, segmentation of a region of interest 35, creation of an isosurface of the region of interest 37a, and generation of a wireframe model of the isosurface 38a may be effected in the general manner described in U.S. Patent No. 5,782,762, which is incorporated herein by reference or any comparable methodologies and algorithms. Alternatively, instead of generating a wireframe model from an isosurface of a segmented

region of interest pursuant to steps 35, 37a and 38a, the wireframe model can be generated at step 38b after a surface model is identified at step 37b from the three-dimensional volume formed at step 34.

5 Patient preparation 32 will be dictated by the organ system to be examined. For example, if the patient's colon is to be examined, the patient is prepared using bowel cleansing and insufflation with gas to distend the colon. In one embodiment, a hand-operated air bulb may be used to distend the colon. 10 The colon is inflated via a rectal catheter by squeezing the air bulb to instill puffs of room air into the colon.

Alternatively, the colon may be inflated with carbon dioxide (CO<sub>2</sub>) or other gas, such as nitric oxide (NO), which exhibits a high lipid solubility and a high partial pressure gradient across the colonic mucosa. The combination of properties provided by carbon dioxide results in less uncomfortable abdominal symptoms for the patient following an examination. 15 However, the high lipid solubility and high partial pressure gradient across the mucosa mean that the carbon dioxide is quickly absorbed by the patient. Therefore, the pressure of carbon dioxide within the colon must be continually monitored and maintained at a beneficial level during the course of an examination. Toward that end, an electronic carbon dioxide insufflator, such as those currently used for laparoscopic surgery procedures, is preferably used to 25 instill the carbon dioxide at a constant flow rate (e.g., 1-15 liters/min) until a preset pressure (e.g., 0.04 atm) is obtained. The insufflator then maintains 30

a constant pressure during the course of the examination.

5 The patient may be placed on a low-residue diet for a 24 hour period preceding the collection of image data. In addition, the patient can be given an oral contrast agent (barium-containing or iodine-containing mixture) to become incorporated in the fecal content to readily distinguish the stool from surrounding soft tissue structures, including the colon wall. The  
10 contrast agent may be used in conjunction with a cathartic agent to rid the bowel of the bulk of stool. The opacified stool may then be digitally removed from the CT image data in order to render an unimpeded colon. Without adequate bowel cleansing or the use of  
15 an oral contrast agent, stool and normal colon wall or colon lesions may be indistinguishable in computed tomography or other imaging modalities. For example, the contrast agent may comprise a barium-containing mixture or an iodine-containing compound for  
20 opacifying solid or liquid stool. In the case of rendering a patient's airways, the patient can be administered a bolus of non-ionic intravenous iodinated contrast agent to aid in distinguishing blood vessels from surrounding airways.

25 Once the patient has been prepared, two dimensional, cross-sectional images of the selected structure are acquired at step 33 with the use of a scanner 422, such as a helical computed tomography (CT) scanner or magnetic resonance imaging (MRI)  
30 scanner. The two-dimensional images are arranged in computer memory 421 (i.e., random access memory (RAM)) to create a three-dimensional data volume at step 34. To create isocubic volume elements (i.e., voxels), an

interpolation method, including but not limited to trilinear interpolation, may be applied.

At step 35, a region of interest is segmented from the three-dimensional data volume. The purpose of segmentation is to isolate a region of interest within the three-dimensional data volume prior to three-dimensional rendering. In general, medical image segmentation is complicated by image noise, partial volume effects, and the fact that similar intensity values are shared by different anatomical structures. However, when a thin-walled soft tissue structure encompasses a gas-filled lumen, segmentation of that structure can be effectuated by selecting the gas column as the region of interest. The gas column can be effectively segmented because the boundary between gas and soft tissue is relatively distinct and sharp. Further, the outside surface of the gas column corresponds to the inside surface of the organ of interest. Similarly, when a thin-walled soft tissue structure encompasses a contrast enhanced blood-filled lumen, segmentation of that structure can be effectuated by selecting the contrast-enhanced blood column as the region of interest. Accordingly, a simple thresholding or region growing technique, along with a morphological dilation away from air or blood vessel and towards soft tissue, can be utilized for segmentation purposes.

However, in order to refine and to improve the accuracy of the segmentation process, especially as selected structures have varying characteristics (e.g., changing diameter, wall thickness, and/or X-ray attenuation values), more sophisticated segmentation procedures can be employed. One such procedure

adaptively adjusts the threshold value defining the region of interest to more accurately segment specific sections of the organ during a region growing process as described in U.S. Patent No. 5,920,319 incorporated herein by reference. Alternatively, the accuracy of segmentation can be improved by refining (or deforming) the wireframe model generated in steps 38a and 38b of Fig. 1 using a geometric deformable model (GDM) at step 40b.

Referring again to Fig. 1, when a patient's colon is to be rendered and a contrast agent has been used to opacify stool and/or residual water, the segmentation step 35 includes the step of digitally removing the opacified stool/water from the CT image data in order to render an unimpeded colon. Digital removal of opacified feces is difficult due to the fact that simple digital subtraction leaves behind edge gradients. The appearance of edge gradients is illustrated in Figs. 15-17. Fig. 15 is a schematic representation of a section of image data representing hypothetical CT image data showing the measured X-ray attenuation values associated with a two dimensional array of voxels. Soft tissue is represented by a value of 100, opacified stool by a value of 200, and air by a value of 0. In this simplified model, actual interfaces between different species is presumed to lie on the boundary between adjacent voxels. Accordingly, an interface 600 between soft tissue and opacified stool appears as the boundary between the voxels having a value of 100 and those having a value of 200. Similarly, an interface 601 between opacified stool and air appears as the boundary between the

voxels having a value of 200 and those having a value of 0.

5 In reality however, sections of actual interfaces are likely to fall within certain voxels. The X-ray attenuation values measured for those voxels will be an average of the X-ray attenuations for the different species comprising the interface. Such volume averaging effects blur the interfaces. As illustrated in Fig. 16, the actual soft tissue/opacified stool boundary falls somewhere within the voxels having a measured value between 100 and 200. Likewise, the actual opacified stool/air interface lies somewhere within the voxels having a measured value between 200 and 0.

10 The effect of a simple digital removal scheme, whereby all values that are 150 or greater are set to zero, on the section illustrated in Fig. 16 is shown in Fig. 17. It is readily apparent that a ridge of voxels having a measured value of 90 is left behind and, therefore, will incorrectly be considered part of the soft tissue. In addition, although the true soft tissue/opacified stool interface lies somewhere within the voxels having a value of 120, the removal scheme automatically assumes that all of those voxels belong to the soft tissue. It will of course be appreciated that the edge effects encountered in real image data will be even more complex, due at least in part to inherent fluctuations in the measured values.

25 In view of the foregoing, when opacified stool is to be subtracted from the image data, an adaptive thresholding technique (as previously described in U.S. Patent No. 5,920,319) and/or a geometrically deformable model approach can be used to model both

the gas column and the adjacent stool. These 3D models can then be analyzed separately, or merged to create a contiguous colon model. For example, the geometric deformable models initialized separately in the gas column and adjacent stool can be allowed to evolve until the boundaries of the two models merge.

Returning to Fig. 1, once the region of interest has been segmented at step 35, an isosurface of the region of interest is created at step 37a. The isosurface can be generated using a variant of a marching cubes algorithm. The isosurface is used to generate a wireframe model at step 38a. The wireframe model comprises a polygonal mesh that corresponds to the surface of the region of interest. Specifically, a plurality of vertices, each vertex having a coordinate position, are interconnected to produce the wireframe model.

Alternatively, a waltzing cubes algorithm can be employed as shown in Fig. 12 to first identify a surface model at step 37b and then to generate a wireframe of the surface model at step 38b without the help of initial segmentation 35. The generated surface can be an isosurface, such as that produced by a marching cubes variant, or it can be a variable-thresholded surface.

First, a voxel near the surface of interest within the volume of data is selected at step 500. This selection can be automated or explicitly identified by the user. Using this selected voxel, the first seed cell and its coordinate are identified at step 502. A cell is defined as the junction of eight adjacent image voxels, four in an upper image plane and four in a lower image plane, such that the eight



voxels form a 2x2x2 cubic cell. A cell coordinate defines its exact location within the volume of data. The first seed cell is defined as any cell near the selected voxel that produces a surface patch belonging to the surface of interest. The first seed cell can be determined by iteratively examining all cells within an increasing radius centered on the selected voxel. Visually, each successive iteration examines all cells that lie on the surface of a sphere centered on the selected voxel as the radius is increased, until the nearest surface patch producing cell is identified. This method is accurate but computationally slow. Alternatively, cells that lie along each direction of each coordinate axis (i.e. -X, X, -Y, Y, -Z, Z) centered on the selected voxel can be examined until the nearest seed cell is identified. This method is fast but requires a more exactly selected voxel.

After the first seed cell and its coordinate are identified, the coordinate is inserted into the waltzing cubes seed queue at step 504. A seed queue stores coordinates of seed cells that are added and removed in subsequent steps.

At step 506, the seed queue is queried to see if it is empty. If the seed queue is empty, the waltzing cubes algorithm is terminated at step 520, since all cells that may produce a connected surface patch of the surface of interest have been analyzed and processed. If it is not empty, then the algorithm proceeds to the next step, and the main loop of the algorithm is entered, because all of the cells which may produce a connected surface patch have not been analyzed. Note that at the start of the algorithm,

the only seed coordinate in the seed queue is the first seed cell coordinate added at step 504.

At step 508, a seed coordinate is removed from the seed queue. At step 510, using the seed cell coordinate, the waltzing cubes cell tree is queried for the corresponding seed cell. The cell tree identifies, organizes and stores all cells within the volume of data that belong to the surface of interest. Additionally, the cell tree maintains cell-to-cell vertex coherency data to ensure that exactly one calculation is made for each shared vertex that lies along the edge between surface patches in two adjacent cells.

The cell tree is structured to contain a set of buckets. A bucket is defined as a linear array of cells where subsequently added cells are appended at the end of the array. A bucket search for a particular cell is done in reverse order, starting with the cell at the end of the array and working back to the beginning. Alternative search algorithms, such as binary search, can be used. However, the reverse order linear search is optimal. Each bucket in the cell tree represents all examined cells within a subset of the volume, called a cell block. The volume is divided into cell blocks of optimal size, the size depending upon the ratio of surface patch producing cells to the total number of cells within the block. An increase in the volume size results in an increase in the number of blocks because the block size remains the same. Typically, a cell block size of 16x16x16 (4096 cells) is near optimal.

At step 512, if the seed cell exists in the cell tree, it has been previously examined, so it is

ignored, and the loop continues by returning to step 506. If the seed cell does not exist in the cell tree, it has not yet been examined and processed, so it is inserted into the cell tree at step 514.

5           At step 516, a surface patch comprised of triangle primitives for the seed cell is produced. The surface patch can be generated by using the seed cell configuration and the corresponding triangulation table for that configuration. The triangulation can be an isosurface patch that has been pre-constructed for each cell configuration and stored in a triangulation table. Alternatively, the triangulation can be a variable-threshold surface patch that is created dynamically using local thresholds, adaptive thresholding or other criteria. The triangulation process uses the cell-to-cell vertex coherency data to reuse vertices computed by other cells and to create new vertices that are not yet computed. The triangulation may produce surface patches that are contained entirely within the seed cell or that span the seed cell and its adjacent cells in the cell tree.

10           At step 518, the adjacent cell coordinates are determined for the seed cell using the seed cell configuration and the corresponding adjacency table for that configuration. An adjacent cell coordinate is the cell coordinate for the corresponding cell that lies adjacent to the seed cell. Each seed cell has up to 6 adjacent cells, and therefore, up to 6 adjacent cell coordinates. An adjacency table contains, for each seed cell configuration, a pre-analyzed list of adjacent cell coordinates that identify cells that each produce a surface patch that is connected to the surface patch produced by the seed cell. Each of

these adjacent cell coordinates is used to query the cell tree to determine whether it has already been examined. If it has been examined, the adjacent cell coordinate is discarded. If it has not been examined, the adjacent cell coordinate is inserted into the seed queue. The algorithm loops back to step 506. When the algorithm terminates at step 520, a wireframe model of a complete surface is produced.

The waltzing cubes method enables the run-time memory required to create the surface of interest to be significantly reduced. Specifically, the memory required is directly proportional to the number of cells that must be analyzed. Since the waltzing cubes method divides the volume of data into cell blocks, entire buckets of cells may be pruned (deleted) once it is known that the cell data is no longer needed. Accordingly, since only those cells which produce surface patches that are a part of the surface of interest are examined, the waltzing cubes method typically examines a small fraction of the total number of cells within the volume, resulting in an algorithm that is fast and efficient.

After the wireframe model has been generated at step 38a or 38b of Fig. 1, connectivity matrices are determined at step 39. The connectivity matrices are data structures which provide information regarding the connectivity between the vertices and polygons which comprise the wireframe model. The connectivity matrices are determined by traversing the polygon and vertex lists associated with the wireframe model, generating sets of immediate neighbor vertices and triangles associated with each vertex and polygon in the lists. For example, the vertex to polygon

connectivity matrix contains, for each vertex in the wireframe model, a list of all polygons that contain that vertex. Likewise, a polygon to polygon matrix lists all adjacent polygons for each polygon in the model.

Using these connectivity matrices, the polygon list of the wireframe model is then re-ordered. The polygons generated using the variant of marching cubes are usually added to, and as such ordered in, the polygon list as the three dimensional volume is traversed. Accordingly, all of the polygons between the first and second images are generated first, followed by the polygons between the second and third images, and so forth. This polygon ordering is not intuitive, nor ideal for analysis or manipulation of the geometry at the local connectivity level. Consequently, in order to group vertices into populations and to analyze the populations, it is advantageous to re-order the polygons into the sequence by which they would be encountered while "traveling" through the lumen of the wireframe model. In addition, the re-ordering facilitates rapid three-dimensional renderings, since hidden objects or objects outside of the field-of-view can be easily identified and removed, thereby reducing the number of polygons rendered and thus increasing the rendering speed.

After connectivity matrices are computed, a normal vector is calculated for each vertex in the wireframe model at step 40a of Fig. 1. The direction of each normal vector is perpendicular to a plane that is tangent to the isosurface at each such vertex, typically pointing away from the object or away from

the lumen of a body organ. The normal vectors at the respective vertices can be computed as the average of the normal vectors associated with each polygon connected to that vertex. The normal vector associated with a polygon is calculated using a vector cross product.

The wireframe model is refined using a geometric deformable model at step 40b. In a geometric deformable model, a surface is treated as a wave front that propagates in space over time until a targeted boundary is reached. With respect to the wireframe model, each vertex of the wireframe model is moved along its normal vector direction at a speed that is a function of both the surface curvature and the doubt (or confidence) level of a boundary estimate at that point. The surface evolves under this type of motion until the targeted boundary is reached, at which time the speed of motion approaches zero. Formally, if  $S(0)$  is an initial smooth surface and  $S(t)$  is a one-parameter family of surfaces generated by wave front propagation, the surface  $S$  should satisfy the following flow equation:

$$\frac{\delta S}{\delta t} = (\phi H - \nabla \phi \cdot \vec{N}) \vec{N} \quad (1)$$

where  $H$  denotes the mean curvature,  $\vec{N}$  is a unit normal vector, and  $\phi$  is a positive differentiable function defined in three-dimensional space that acts as a stopping function (i.e.,  $\phi$  approaches zero as the surface nears the targeted boundary).

A direct solution for this equation is difficult because of the topology changes and singularities that are created during the course of surface evolution. Accordingly, special techniques are typically used to facilitate convergence of Eqn. (1). For example, a level-set technique can be used, such as the one proposed by Osher and Sethian in "Fronts propagation with curvature dependent speed: algorithms based on Hamilton-Jacobbi formulations", J. Computational Physics, 1988, 79: 12-49, which is hereby incorporated by reference. In the level-set approach, the surface S is embedded as a level surface  $\Psi=0$  in a three-dimensional scalar function  $\Psi$ . The evolving family of surfaces  $S(t)$  is then generated from the level surface of an evolving family of scalar functions  $\Psi(t)$ . As shown in Fig. 3, evolving contours are treated as zero-level curves of the evolving function F. By shifting the scalar function F downward from time  $t_1$  to time  $t_2$  and extracting the zero level curves, the net effect of shrinking the circular contour (Fig. 3(a)) is achieved (Fig. 3(b)). If F is shifted down even further, the circular contour will appear to split into two smaller segments.

For the embedded level-surface to evolve according to Eqn (1), the scalar function  $\Psi$  should evolve according to the equation:

$$\frac{\delta \Psi}{\delta t} = \phi \|\nabla \Psi\| \left( \operatorname{div} \left( \frac{\nabla \Psi}{\|\nabla \Psi\|} \right) + v \right) + \nabla \phi \cdot \nabla \Psi. \quad (2)$$

The divergence operator  $\text{div}\left(\frac{\nabla\Psi}{\|\nabla\Psi\|}\right)$  computes the mean

curvature  $H$  of the embedded level surface from the scalar function  $\Psi$ . The constant,  $v$ , determines if the surface evolves inwardly or outwardly. The stopping function,  $\phi$ , can be defined as:

$$\phi = \frac{1}{1 + \|\nabla G_{\sigma} * I\|}$$

where  $G_{\sigma}$  is a Gaussian filter whose characteristic width is  $\sigma$  and  $*$  denotes a convolution operator. Images convoluted with the Gaussian filter before gradient computation are smoother and produce more accurate results. The purpose of the Gaussian filter is to smoothly force the evolution speed to zero at points where high intensity gradients exist.

More generally, the stopping function can be characterized as a doubt function that approaches 0 when there is no doubt that the wave has reached a boundary and 1 otherwise. Using this characterization better stopping functions can be designed so that they incorporate not only edge gradient information, as in Eqn(2), but also other critical information such as noise model and scale level. Furthermore, Bayesian statistical models can be used to properly define the doubt function since many factors contribute to one's doubt (or confidence) about any point being on a boundary.

Traditionally, a major problem associated with



the use of geometric deformable models has been poor computational efficiency due to the fact that multiple iterations are required at numerous grid points. One approach to improve the computational efficiency has been the use of narrow-band or tube methods as described by Malladi et al. in "Shape modeling with front propagation: a level set approach", IEEE Trans. Pattern Analysis and Machine Intelligence, 1995, 17:158-175, which is hereby incorporated by reference. Although useful in some applications, the narrow-band method is inefficient when dealing with large wireframe models, such as those representing a colon. Accordingly, special constraints may need to be added to reduce the processing time for large objects. In one embodiment, a three-dimensional region-growing model is used as a starting point for the deformable model approach. Alternatively, the deformable model approach may be applied to a low resolution dataset to produce an intermediate wireframe model. The intermediate wireframe model is then translated to a higher resolution dataset. Further, since the original volume of data typically has a high spatial resolution, the deformable model approach can be applied to a subsampled volume (e.g., one-eighth of the original volume of data) to produce an intermediate wireframe model without significantly affecting the shape of the rendered image. The intermediate wireframe model is then applied as a starting point for processing of the complete volume of data.

The vertices of the wireframe model can be analyzed and grouped into populations having abnormal wall structure as shown at steps 41-45 in Fig. 1.

The wall thickness of the structure at each vertex is measured at step 41. The wall thickness at a given vertex can be determined by measuring the distance from the inner surface of the selected object to its outer surface or some other position between the inner and outer surfaces. For body parts such as the colon or airways, the selected structure's wall is composed of soft tissue and the lumen of the structure comprises gas. Accordingly, the inner surface corresponds to a gas/soft tissue interface. Further, the colon and airways are typically surrounded by fat, gas, or other non-soft tissue material (i.e., contrast-enhanced blood vessels, bone, or calcification). Accordingly, the outer surface corresponds to the soft tissue/gas, soft tissue/bone, soft tissue/fat, or other such interface. The X-ray attenuation factors for room air (approximately -1024 HU to -425 HU), fat (about -200 HU to -20 HU), and other non-soft tissue structures such as contrast-enhanced blood vessels and bone (approximately >100 HU) are distinct from the soft tissue component (about 20 HU to 100 HU) of the airways or colon wall. Accordingly, the wall thickness at a vertex on the wireframe model can be calculated from the volume data of the selected organ, which can be measured at regularly spaced intervals along each normal vector. Alternatively, wall thickness can be estimated by analyzing differential characteristics of the intensity profile along the normal direction. For example, the transition from gas-filled lumen to wall tissue and then from wall tissue to fat or other software tissue can be captured by finding the first two peaks in the gradient of the intensity profile.

Referring to Fig. 6, a wall thickness at a vertex can be calculated by analyzing the intensity profile along the normal vector originated at the vertex. An intensity profile of a vector is defined as an array of interpolated x-ray attenuation factors (voxel values) along regularly spaced intervals along a vector originating at the vertex. The analysis of the intensity profile may include several heuristics depending upon the intensity profile type. One simple heuristic is to measure the thickness as the distance between the vertex and the point at which the attenuation factor drops below that of air (-425 HU) as seen in Fig. 6a. Another heuristic is to measure the thickness as the distance between the vertex and the point at which the attenuation factor exceeds above that of soft tissue (60 HU) as seen in Fig. 6b. Yet another heuristic is to measure the thickness as the distance between the vertex and the point at which there is a local minimum as seen in Fig. 6c. These heuristics are often too basic and are insufficient to accurately determine complex intensity profiles such as that in Fig. 6d. Therefore, a more mathematical analysis of the intensity profile is required for accurate wall thickness calculation. The advanced analysis include differential characteristics of the intensity profile, discrete first/second/third degree gradient profiles, polynomial curve fitting along the intensity profile and solutions to differential equations based on the curve, and other knowledge based methodology. For optimal wall thickness measurement, a weighted combination of basic and advanced analysis is employed, and the results are averaged and normalized to reduce inaccuracies.

Alternatively, the wall thickness at the vertex on the wireframe model can be calculated from the volume of data of the selected organ near the vertex. For example, another approach to estimate wall thickness is to classify the voxels in the vicinity of the segmented lumen surface into a small number of intensity groups. The shape of the resulting groups can be analyzed to determine the voxels that belong to wall and the wall thickness can then be estimated from these voxels.

Since the wall thickness of body structures such as the colon may be inadvertently or artificially increased, such as during musculature contraction of the colon at certain positions along the colon structure, many of the regions of abnormal thickness may not be indicative of true lesions. Instead, the thickened regions may represent normal structural elements, such as regions where the colon is contracted or where it touches solid organs. Accordingly, it can be beneficial to further analyze and characterize each vertex associated with abnormal thickness to refine the list of potentially abnormal vertices. Additional local shape features, computed at step 42, can be used to characterize and associate vertices as being abnormal.

For example, a local convexity value at each of the surface vertices can be determined at step 42. The local convexity values can be determined for all vertices, or for only those vertices associated with abnormal wall thickness. For each vertex, its neighboring vertices within a connected neighborhood of a pre-determined size (e.g., 3), is determined using the connectivity matrices. A local convexity

value for each vertex,  $v$ , is computed as:

$$\bar{C} = \sum_{i=1}^n D(v_i)$$

5

wherein  $D(v_i)$  is the distance from the  $i^{\text{th}}$  neighboring vertex to a plane through  $v$  and perpendicular to  $v$ 's normal vector, and  $n$  is the number of neighboring vertices. Vertices associated with a convexity value of greater than a pre-defined maximum (e.g.,  $>2$ ) can be considered abnormal.

10

In addition, the local curvature at each vertex in a population can also be determined at step 42. The curvature reflects another shape feature of the object's surface at a given vertex. Vertices associated with broad curvatures indicate regions that are relatively flat. Accordingly, if the user wishes to identify only regions having thicknesses that vary significantly over short distances, those vertices with small curvatures may be labeled as abnormal.

15

20

The local curvature at each vertex in a population can be determined by summing the scalar distance between each of a user-determined number of levels,  $N$ , of adjacent vertices and the plane perpendicular to the normal of the vertex. The result is then normalized by the surface area outlined by the  $N^{\text{th}}$  level of adjacent vertices. Because the direction of the curvature is ignored, the technique is able to detect areas of small curvature.

25

Alternatively, local curvature can be computed at each vertex from the wireframe model represented as a level surface. In particular, as shown in Fig. 13, the mean curvature at each vertex of the wireframe model is determined at step 550. The mean curvature  $H$  (i.e., the average of the two principal curvatures) of the level surface  $\Psi$  is given by:

$$H = \frac{\Psi_{xx}(\Psi_x^2 + \Psi_z^2) + \Psi_{yy}(\Psi_x^2 + \Psi_z^2) + \Psi_{zz}(\Psi_x^2 + \Psi_z^2) - 2\Psi_x\Psi_y\Psi_{xy} - 2\Psi_y\Psi_z\Psi_{yz} - 2\Psi_x\Psi_z\Psi_{xz}}{2(\Psi_x^2 + \Psi_y^2 + \Psi_z^2)^{3/2}}$$

The Gaussian curvature  $K$  is similarly determined according to the equation at step 552:

$$K = \frac{\left| \frac{\Psi_x^2}{\Psi_{xx}\Psi_{yy} - \Psi_{xy}^2} + \frac{\Psi_y^2}{\Psi_{xx}\Psi_{zz} - \Psi_{xz}^2} + \frac{\Psi_z^2}{\Psi_{yy}\Psi_{zz} - \Psi_{yz}^2} + 2\frac{\Psi_x\Psi_y}{\Psi_{xx}\Psi_{zz} - \Psi_{xz}^2} + 2\frac{\Psi_x\Psi_z}{\Psi_{xx}\Psi_{yy} - \Psi_{xy}^2} + 2\frac{\Psi_y\Psi_z}{\Psi_{yy}\Psi_{zz} - \Psi_{yz}^2} \right|}{2(\Psi_x^2 + \Psi_y^2 + \Psi_z^2)^{3/2}}$$

Each vertex of the wireframe model can be classified as one of four types at step 554. The point is classified as elliptic if  $K(p) > 0$ , as hyperbolic if  $K(p) < 0$ , as parabolic if  $K(p) = 0$  and  $H(p) \neq 0$ , or as planar if  $K(p) = 0$  and  $H(p) = 0$ . This classification scheme enables the vertices to be grouped into patches having vertices having the same characteristic shape at step 556. Since polyps protrude into the colon lumen, only elliptic patches are labeled as representing likely polyps at step 558.

Additionally, elliptic patches that point outwardly from the colon are eliminated at step 560 using the signs of the two principle curvatures,  $k_1$  and

$k_2$ . The principal curvatures,  $k_1$  and  $k_2$ , are derived from the mean and Gaussian curvatures to be

$$k_1 = H + \sqrt{H^2 - K} \quad \text{and} \quad k_2 = H - \sqrt{H^2 - K}. \quad \text{Only when } k_1 \text{ and } k_2$$

are both negative (or positive depending on coordinate system used) does the elliptic patch represent a potential lesion pointing inward with respect to the lumen.

At step 44, vertices representing an abnormality are selected. Those vertices meeting selected criteria can be identified as abnormal vertices. For example, vertices having abnormal thickness and/or convexity and/or curvature criteria may be identified as abnormal vertices. In a specific application, the wall thickness values, local convexity values, and local curvature values can be used, independently or in conjunction with each other, to either accept or reject vertices as being abnormal. The individual parameters can be combined using a logical AND operation.

At step 45, the vertices on the wireframe model associated with abnormal structure (i.e., having a combination of associated abnormal wall thickness and/or abnormal shape, such as abnormal local convexity and/or abnormal local curvature) are grouped into populations. The re-ordered connectivity matrices are used to determine if vertices associated with abnormal parameters are directly connected to other vertices that are also associated with abnormal parameters. Accordingly, each formed population represents a potential abnormal lesion. In addition, each population can be further analyzed and characterized by its size, dimensions, or other

statistical quantities.

A cross sectional area (CSA) of surface model is defined as the area outlined by the contour formed by the interaction of the surface model 900 with an arbitrary CSA plane 902 as shown in Fig. 8. The cross sectional area at the site of a lesion may be used to calculate the degree of stenosis, or narrowing, caused by the lesion at that location. The cross sectional area may also be used with a central path algorithm to determine areas of stenosis in a lumen of an organ. The surface model is represented as a wireframe mesh of triangles. Using the connectivity matrices, all triangles in the wireframe which intersect the CSA plane are identified as shown in Fig. 9a. For each triangle, edges of the triangle which intersects the CSA plane are identified, and intersecting vertex point between each edge and the CSA plane are computed. The set of these intersecting vertex points represents an approximation of the cross sectional area which is to be computed, and it is defined as the CSA contour. The CSA contour lies completely in the CSA plane as shown in Fig. 9b. The CSA contour vertices are used to triangulate a set of adjacent triangles that span the entire contour. The triangulation may simply define the edges of each triangle as consecutive contour vertices as shown in Fig. 9c. Alternatively, a centroid point for the contour may be computed and used for a more accurate triangulation as shown in Fig. 9d. Because the generated triangles span the contour, the cross sectional area of the contour is simply defined as the sum of the areas of each triangle within the contour. This method is a numeric approximation that is



sufficiently close to the true cross sectional area.

Each population is further analyzed according to aggregate shape features at step 46 to reduce the occurrence of false positive findings. Since the structure of body organs may appear abnormal, either inadvertently or artificially, such as during muscular contraction of the colon at certain positions along the colon's structure, many of the abnormal populations may not be indicative of true lesions.

Instead, these populations may represent normal structural elements, such as regions where the colon is contracted or where it touches solid organs. Accordingly, it may be advantageous to further analyze and characterize each population identified as having an abnormal structure to further refine the list of potential lesions and to increase the likelihood that a detected population represents a true abnormality.

For example, the shapes of the populations can be analyzed, characterized, and accepted or rejected as being abnormal as illustrated in detail in Fig. 2. At step 120 a centroid for each population is calculated. The centroid can be computed as a weighted sum of the positions of each vertex within the population. The individual vertices can be weighted by their associated wall thickness values or other parameters. Once the centroid has been determined for a population, the height and normal vector for that population can be determined.

A population normal vector is calculated for each population at step 121. The population normal vector can be computed as a sum of the normal vectors at each of the vertices within the population, and the individual vertex normal vectors can be weighted by

their corresponding wall thickness values or other parameters.

The convexity value,  $\langle C \rangle$ , for each population is determined at step 122. The convexity is a measure of the direction and magnitude of the shape of the surface of the population. As shown in Fig. 5, the convexity of a population is computed as the sum of the distances 94 from vertices 95 in the population to a plane 96, perpendicular to the population normal vector 97 passing through the centroid 98, according to the equation:

$$C = \sum_{i=1}^n D(v_i)$$

wherein  $D(v_i)$  is the distance from the  $i^{\text{th}}$  vertex of the population to the plane 96 and  $n$  is the number of vertices 95 within a predetermined space relative to the centroid. Since the normals at each vertex are typically assigned to point away from the center of the object, or the lumen of the object, a population which protrudes into the lumen of the object would have a positive convexity value, whereas a population that projects away from the lumen would exhibit a negative convexity value. When the structure is a colon, populations with a negative convexity value are excluded from being considered abnormal, since abnormal colon masses tend to project into the lumen of the colon and would have a positive convexity value.

Further, the higher the magnitude of the convexity value, the greater the slope of the surface of the population. When the structure is a colon,

only populations having a positive convexity value above a minimum value are reasonably expected to be potential lesions, since some cancerous colon masses are generally manifested by steeply sloped growths that protrude into the lumen of the colon.

The height (H) of each population is determined at step 123. The height is calculated as the difference between the distance of the vertex farthest from a plane 96, perpendicular to the population normal 97 and passing through the centroid 98, and the distance of the vertex closest to that plane according to the equation:

$$H = \text{MAX } D(v_i) - \text{MIN } D(v_i).$$

The height provides an indication as to whether a population is reasonably expected to be a true lesion since populations with large heights are more likely to represent abnormal masses.

The principal curvatures,  $k_1$  and  $k_2$ , can also be used to reduce the likelihood of false positive identifications. Generally, polyps are nearly symmetrical and, accordingly, patches representing true polyps are likely to arise when the two principal curvatures on the top of a potential lesion are similar. Further, normal colon haustra have ridge-like shapes and, therefore, potential lesions having principal curvatures,  $k_1$  and  $k_2$ , which are substantially different are eliminated as likely polyps at step 562.

Other shape features may be used to include or exclude populations in the list of potential lesions. For example, if a population has a long axis dimension

that is greater than a multiple (e.g., 10 times) of the short axis dimension, and the short axis dimension is, for example, less than 5 mm wide, then the population might be considered to be normal. Further, a potential lesions "roundness" could be quantified and used to rank potential lesions. Likewise, weighting wall thickness determinations with shape features may provide valuable heuristics for classifying potential lesions. Accordingly, the shape features discussed herein are included only by way of example, and are not intended to limit the scope of the invention.

Referring again to Fig. 1, after the populations with abnormal wall structure (i.e., wall thickness, convexity, or other shape features) have been identified and characterized, populations that fail to meet certain criteria are eliminated as potential lesions at step 48.

At step 50, those populations which are still considered to represent potential lesions are sorted and arranged in an electronic listing. The listing can be sorted according to such properties as size, convexity, curvature, height, mean wall thickness, standard deviation of wall thickness, other statistical quantities, or a combination of such parameters (e.g., the product of the group convexity value, C, and the group height value, H). Each population in the listing could be linked to a three-dimensional camera position that best illustrates it in the three-dimensional display. Accordingly, the user can choose a population from the listing and thereby be presented with a view of the abnormal area as it appears in the three-dimensional rendering. One

means of linking the population to a three-dimensional rendering camera position is by providing the user with a three-dimensional camera view of the population that is parallel to the population normal and centered on the centroid of the population, and set at a chosen distance from the centroid. In one embodiment, the user is presented with two views of the selected population, one from an external perspective and the other from an internal perspective (i.e., from within the lumen of the structure). Lesion locations, either identified manually by a user or by the above algorithms can also be used to generate lesion maps, i.e., two dimensional or three dimensional renderings of entire colon with lesion sites marked.

A three-dimensional rendering of the wireframe model is displayed at step 51. In one embodiment, the user can navigate through a three-dimensional rendering of the wireframe model which is displayed on the monitor 28. The detected abnormal populations are displayed with a contrasting color scheme to distinguish them from the remainder of the rendered structure, so that the abnormal populations can be quickly and easily identified by human visual inspection. Further, the color scheme can reflect variations in wall thickness, convexity, or curvature, or some other parameter. For example, the normal portions of the rendered structure can be displayed in shades of pink while the potentially abnormal populations are displayed in a rainbow of colors. More specifically, vertices could be assigned colors ranging from dark blue (for vertices with minimum abnormal thickness) to bright blue (for vertices with maximum abnormal wall thickness) to provide a stark

contrast to the pinkish color assigned to the normal portions of the rendered structure.

In an alternate embodiment, the user is presented with a volume rendered display centered on the centroid of a selected abnormal population. For example, a volume-rendered view of a subvolume of user defined size centered on the centroid of the population can be displayed for visual inspection. Volume rendering is a visualization technique that creates a three-dimensional image using all of the volume data, rather than only an extracted surface representation. Traditionally, volume rendering is accomplished by projecting a series of rays from the user's viewpoint through the data volume. The color at each point (i.e., pixel) in the resulting three-dimensional image is determined by calculating a weighted sum of all voxels along each ray. This weighting factor is determined by a transfer function that is computed during a process called classification. Additionally, voxels belonging to a lesion may be selectively enhanced with color and/or opacity values.

Altering the transfer function during volume rendering can reveal different characteristics of the rendered data and, thus, it is often desirable to interactively reclassify and re-render the data. However, the rate at which images are rendered can be slow for data volumes as large as the ones typically used in medical imaging. Accordingly, it can be beneficial to use texture mapping to dramatically increase the rate at which volume rendering is performed. In texture mapping, volume data is considered to be a series of two-dimensional textures

that are interpolated, projected, and displayed rapidly on a series of polygons using specialized computer hardware.

5 The texture mapped volume rendering method uses high-speed texture memory and blending hardware available on a Silicon Graphics' computer workstation to approximate the results of the more computationally expensive ray-casting algorithm. The volume data is first loaded into texture memory. The textures are  
10 then semi-transparently mapped onto a series of parallel two-dimensional rectangles that correspond to slices of the volume. The texture mapped rectangles are then drawn and blended to allow the user to see through the volume. The colors and transparencies of the resulting three-dimensional object are controlled by color and opacity transfer functions. The entire volume can then be viewed by stacking the texture mapped rectangles in the proper order and looking  
15 toward the stack. Since viewing the stack along one of its edges does not optimally display the volume, a new stack is generated which is orthogonal to the original stack. This new stack is then used to display the volume along an edge. Likewise, for any additional viewing angles, additional stacks may be  
20 required. Generating the stacks is a dynamic process and each stack can be generated on an as needed basis. Texture mapping can use standard two-dimensional texture memory or faster three-dimensional texture memory.

30 In yet another alternate embodiment, upon selection of a population from the listing, the user can be presented with a three-dimensional, surface-rendered wireframe model of the population embedded

within a volume-rendered display using texture memory techniques. Such a display is useful to show the three-dimensional relationships between the wireframe model and the volume data. Embedding a surface-rendered model inside a volume rendering can be accomplished using texture memory since both techniques use semi-transparent or opaque surfaces to generate their imagery. However, if the surface rendering is semi-transparent, care must be taken to ensure that all objects are blended in the proper order (typically, from back to front). All or part of the volume can be rendered. For example, to view a small structure using volume rendering, a subvolume of data surrounding that structure can be rendered, as opposed to displaying the entire volume of data. The point about which the subvolume rendering is performed can be selected manually by the user, or automatically by the above algorithms. It should also be appreciated that the multiplanar views and volume rendering can be combined in a similar manner.

If the user wants to view the results of the various rendering techniques using separate display windows, the user can clone the display window 200 so that the two (or more) display windows share the same data. However, each display window could render using different techniques (i.e., multiplanar display, surface rendering, volume rendering). Accordingly, the user could open a single window with all three rendering techniques and then clone the window three times to view each technique separately. Changes made to the data in any one of the connected windows would be propagated to all the cloned windows. Camera orientations could be synchronized between the



5 separate windows to lock the views together. Position tracking techniques could show the camera position of one viewer in another, so that the user could follow the path of a three-dimensional camera moving inside the structure in one display window from an external perspective using a cloned display window with another three-dimensional camera from an overhead position.

10 An example of a computer console display 100 useful for displaying a three-dimensional rendering of a structure is shown in Fig. 7. The display 100 comprises a first window 101 for displaying a volume rendered image of the structure and a second window 102 for displaying a surface rendered image of the structure. Alternatively, the second window 102 can display a surface rendered image of the structure embedded within a volume rendered display, as described above. In addition, three multiplanar windows, 103, 104, and 105, are provided in a third window for displaying axial, coronal, sagittal, or oblique two-dimensional slices through the structure at a user-defined point, or at the centroid of a chosen population.

15  
20  
25  
30 Alternatively a single window display 200, as shown in Fig. 10, can be used to display a three-dimensional rendering of the structure. The single window display comprises a single three-dimensional display window for presenting a combination of multiplanar images 203, 204 and 205 with a volume-rendered image 207 centered on a point located within a surface-rendered image 209. The display presents a holistic view of all the data that can aid in the visualization and understanding of the inter-relationships of the various rendering techniques

(i.e., surface rendering, volume rendering, and intersecting two-dimensional planes). The various rendering techniques can exist together because they are all based on the same patient coordinate system supplied by the image scanner, and they are all rendered using shared three-dimensional graphics techniques (texture memory) and a common graphics library. The intersecting planes can be rendered as opaque textures (as opposed to semi-transparent textures used in volume rendering) mapped onto two-dimensional rectangles using texture mapping and drawn in their proper orientation. The user can interactively scroll through slices parallel to any of the intersecting planes and can view any combination of planes (e.g., axial and/or sagittal and/or coronal planes), or the user can create and view an oblique slice plane. The planes can be rendered together with the surface data to show the exact relationship between the wireframe model and the volume (by viewing the intersections and orientations of the surface rendered wireframe model and the two-dimensional slices). They can be viewed together because the planes are rendered as any other kind of surface (i.e., a flat, two-dimensional rectangle).

The method and system of the present invention can also be used to guide needle biopsy procedures of diagnosed lesions as illustrated in Fig. 14. One example is biopsy guidance of cancerous lymph nodes in patients with lung cancer via transbronchial needle aspirations (TBNA). At steps 800 and 802, the patient's airway, lymph nodes, blood vessels, and any cancerous masses are segmented and rendered in a three-dimensional display. The rendering is performed

as described above. Of course, the structures must be accurately rendered in order to provide a reasonable likelihood that the procedure will be successful. An alternative approach to automatically rendering segmenting the lymph nodes and/or the cancerous masses from the volume data, spherical "targets" can be manually placed within lesions at lesion locations previously identified on the multiplanar data images to indicate the position and approximate size and shape of the targeted object. Alternatively, irregular targets can be modeled by having the user free-hand draw regions of interest on the multi-planar images from which three-dimensional targets can be rendered.

Once the lymph nodes and cancerous masses have been identified and rendered, each lymph node and cancerous mass which is to be biopsied is labeled as a "targeted structure" at step 804. A zone, or striking distance, is then defined around each of the targeted structures in step 806. The zone represents a volume of space surrounding the targeted structure which is sufficiently close to the structure to enable a biopsy needle to reach the targeted structure and biopsy tissue from the targeted structure. At step 808, blood vessels and other mediastinal tissues may be rendered. Rendering of the mediastinal tissues is performed because it is important to avoid puncturing major blood vessels during the needle biopsy procedure.

An endoscopic biopsy procedure, such as TBNA, is performed with the actual biopsy needle retracted within the instrument channel of an endoscope while the endoscope is inserted in the airway. As soon as the endoscope tip is within striking distance of a

pre-selected target, the needle is extended into the target and then retracted back into the endoscope's instrument channel, hopefully containing a sample of tissue. Unfortunately, doctors often make "blind" biopsies into tissue since they cannot visually see through an opaque organ, hence the purpose of computer-simulated guidance. "Striking distance" is defined as the maximum distance from the target that the endoscope tip can located in order to obtain an adequate sample when the needle is extended. In clinical practice, this distance should be at most half the length of the needle-- $\frac{1}{2}$  cm for a 1 cm needle--so that when the needle is extended a sample of at least  $\frac{1}{2}$  the needle length is obtained. Also, both the scope's position and orientation must be taken into account to decide if the needle will be able to effectively biopsy the target, otherwise, if the scope tip is within striking distance but pointed away from the target, then the extended needle will not hit the target. If the tip is not pointed directly at the target but is pointed at an oblique angle to the target, the needle may glance off the side of the target and not obtain an adequate tissue sample. And if the target's diameter is smaller than the needle length, there is a chance that the tip may completely penetrate the target and biopsy tissue on the other side of the target.

The simulated endoscope containing a biopsy needle within its instrument tract is then inserted into and moved within the patient's airway at step 810. As the simulated endoscope moves through the airway, the position of the tip of the needle is monitored at step 812. The monitoring can be done

either on a continuous basis or intermittently. If desired, the movement of the simulated endoscope tip through the airway can be recorded and documented at step 811.

5           Once the position of the simulated endoscope tip with its needle is determined, a rendered image of the needle tip within the rendered image is displayed at step 814. Accordingly, the physician performing the biopsy procedure can visually monitor the movement of  
10           the simulated needle through the airway in relation to surrounding anatomy. In order to more realistically simulate the view that would be obtained using a conventional bronchoscope, the displayed image can be altered to reflect the distorted camera view of  
15           conventional bronchoscopes. In addition to displaying a rendered image of the simulated needle tip within the airway, an external view of the airway showing the position and orientation of the needle can be simultaneously displayed.

20           At step 816, it is determined whether the needle tip is within the zone, or within striking distance, of a targeted structure. If the needle tip is not within the striking distance of a targeted structure, then the distance and trajectory for the needle to  
25           reach a targeted structure is determined at step 818. The needle is then repositioned within the airway at step 810 using the calculated trajectory and distance as a guide for the movement of the needle.

30           However, if the simulated needle tip is within the striking distance of a targeted structure at step 816, then a ready-to-strike target proximity indicator is issued at step 820. The indicator serves to signal the physician to biopsy the targeted structure, at

step 822. There are four types of target proximity indicators: (1) "getting close" indicator that increases in intensity as the endoscope tip approaches the target, (2) "strike zone" indicator that tells when the endoscope tip is within striking distance of the target and in the proper orientation to obtain an adequate tissue sample, (3) "too close" indicator that signals when the scope tip is too close to the target, and (4) "close, but no cigar" indicator that tells when tip is close enough to strike but is not pointing in the right direction for obtaining a good sample. In addition to these four indicators, a fifth type or alarm can be activated when the needle is in danger of penetrating vital organs near the target and putting the patient at risk.

It will be recognized by those skilled in the art that changes or modifications may be made to the above-described embodiments without departing from the broad inventive concepts of the invention. It should therefore be understood that this invention is not limited to the particular embodiments described herein, but is intended to include all changes and modifications that are within the scope and spirit of the invention as set forth in the claims.

## WHAT IS CLAIMED IS:

1. A system for interactively displaying three dimensional structures comprising:
  - a. volume formation means for forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with a three dimensional body;
  - b. segmentation means for segmenting a region of interest from the volume of data based on selected values of the physical property representing the region of interest;
  - c. means for producing a wireframe model of the segmented region of interest, the wireframe model comprising a plurality of vertices, each vertex having a coordinate position;
  - d. means for refining the wireframe model by adjusting the coordinate positions of the vertices to more accurately represent the region of interest; and
  - e. rendering means for rendering the wireframe model in an interactive three-dimensional display, therein producing a virtual three-dimensional environment.
2. The system of Claim 1 wherein the segmentation means approximates the region of interest and the refinement means adjusts the coordinate positions of the vertices along a normal vector associated with each vertex.
3. The system of Claim 2 wherein the refinement means adjusts the coordinate positions of the

vertices along a normal vector associated with each vertex as a function of a surface curvature at the coordinate position.

4. The system of Claim 2 wherein the refinement means adjusts the coordinate positions of the vertices along a normal vector associated with each vertex by determining an inner wall position and an outer wall position from a first and a second degree derivative associated with the coordinate positions of the vertices.
5. The system of Claim 1 wherein the segmentation means, the wireframe producing means, and the refining means are located remotely from the volume formation means.
6. A system for interactively displaying three dimensional structures comprising:
  - a. volume formation means for forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with a three dimensional body;
  - b. segmentation means for segmenting a region of interest from the volume of data based on selected values of the physical property representing the region of interest;
  - c. isosurface creation means for creating an isosurface of the segmented region of interest;
  - d. model means for producing a wireframe model of the segmented region of interest; and



- e. rendering means for rendering the wireframe model in an interactive three-dimensional display, therein producing a virtual three-dimensional environment.
7. The system of Claim 6 wherein the isosurface creation means comprises:
- a. means for identifying a seed voxel from the segmented region of interest; and
  - b. means for producing an isosurface patch associated with the seed voxel.
8. The system of Claim 6 wherein the segmentation means, the isosurface creation means, and the wireframe producing means are located remotely from the volume formation means.
9. A system for interactively displaying a three-dimensional rendering of a structure having a lumen and indicating regions having abnormal structure comprising:
- a. volume formation means for forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with the three-dimensional structure;
  - b. segmentation means for segmenting a region of interest from the volume of data based on a selected value of the physical property representing the region of interest;
  - c. wireframe generation means for generating a wireframe model of the segmented region of interest;

- d. vertex grouping means for grouping the vertices of the wireframe model into regions having abnormal structure;
- e. analyzing means for analyzing the regions having abnormal structure; and
- f. rendering means for rendering the wireframe model in an interactive three-dimensional display to indicate the regions having abnormal structure.
10. The system of Claim 9 wherein the analyzing means comprises means for evaluating two principal curvatures associated with each point in an elliptical region.
11. The system of Claim 10 wherein the analyzing means comprises a means for identifying ridges.
12. The system of Claim 10 wherein the segmentation means, the wireframe generating means, the vertex grouping means, and the analyzing means are located remotely from the volume formation means.
13. A computer-implemented method for interactively displaying a three-dimensional rendering of a structure having a lumen and indicating regions having abnormal structure comprising the steps of:
- a. forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with the three-dimensional structure;

- 52

- g. repeating step 14.c through step 14.f until the seed voxel queue is empty.
15. The method of Claim 13 wherein the segmentation step, the wireframe generating step, the grouping step, and the analyzing step are performed remotely from the volume formation step.
16. A computer-implemented method for interactively displaying a three-dimensional rendering of a structure having a lumen comprising the steps of:
- a. forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with the three-dimensional structure;
  - b. segmenting a region of interest from the volume of data based on a selected value of the physical property representing the region of interest;
  - c. creating a surface of the region of interest;
  - d. generating a wireframe model of the surface;
  - e. deforming the surface to more closely conform to the region of interest; and
  - f. rendering the wireframe model in an interactive three-dimensional display.
17. The method of Claim 16 wherein the segmentation step, the wireframe generating step, and the refining step are performed remotely from the volume formation step.

18. A computer-implemented method for interactively displaying a three-dimensional rendering of a structure having a lumen comprising the steps of:
- a. forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with the three-dimensional structure;
  - b. segmenting a region of interest from the volume of data based on a selected value of the physical property representing the region of interest;
  - c. generating a wireframe model of the surface;
  - d. refining the wireframe model of the segmented region of interest to more accurately represent the region of interest; and
  - e. rendering the wireframe model in an interactive three-dimensional display.
19. The method of Claim 18 wherein the segmentation step approximates the region of interest and the refinement step adjusts the coordinate positions of the vertices along a normal vector associated with each vertex.
20. The method of Claim 19 wherein the refinement step adjusts the coordinate positions of the vertices along a normal vector associated with each vertex at a rate that is a function of a surface curvature and the confidence level of a boundary estimate at the coordinate position.

21. The method of Claim 19 wherein the refinement step adjusts the coordinate positions of the vertices along a normal vector associated with each vertex by determining an inner wall position and an outer wall position from a first and a second degree derivative associated with the coordinate positions of the vertices.
22. The method of Claim 18 wherein the segmentation step, the wireframe generating step, and the refining step are performed remotely from the volume formation step.
23. A method for guiding biopsy of a tissue by accessing the tissue via a three-dimensional structure having a lumen comprising the steps of:
- a. forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with the three-dimensional structure and at least one physical property associated with the tissue;
  - b. segmenting a first region of interest from the volume of data based on a selected value of the physical property associated with the three-dimensional structure;
  - c. creating a first surface of the first region of interest;
  - d. generating a first wireframe model of the surface of the first region of interest;
  - e. segmenting a second region of interest from the volume of data based on a selected value of the physical property associated with the

- tissue;
- f. creating a second surface of the second region of interest;
  - g. generating a second wireframe model of the isosurface of the second region of interest;
  - h. rendering the first and second wireframe models in an interactive three-dimensional environment;
  - i. inserting a simulated biopsy needle within the lumen of the three-dimensional structure;
  - j. determining the position of the simulated biopsy needle within the three-dimensional structure; and
  - k. displaying the rendered, interactive three-dimensional display from a perspective indicating the position of the biopsy needle within the three-dimensional structure.
24. The method of Claim 23 comprising the step of indicating the position of the simulated biopsy needle relative to the position of the tissue.
25. The method of Claim 24 wherein the indicating step comprises the steps of:
- a. generating a zone around the simulated tissue; and
  - b. issuing an indicator when the tip of the simulated biopsy needle is within the zone.
26. The method of Claim 24 wherein the indicating step comprises the step of determining a distance and a trajectory from the simulated biopsy needle

to the simulated tissue.

27. The method of Claim 13, 16, or 18 comprising an image acquisition step wherein a series of two-dimensional images representing at least one physical property associated with the structure is acquired, wherein the image acquisition step comprises a patient preparation step.
28. The method of Claim 27 wherein the patient preparation step comprises the step of administering to a patient a nonionic intravenous iodinated contrast bolus.
29. The method of Claim 27 wherein the patient preparation step comprises the step of administering to a patient an oral contrast agent for opacifying stool and water.
30. A system for interactively displaying three dimensional structures comprising:
  - a. volume formation means for forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with a three dimensional body;
  - b. modeling means for producing a wireframe model of a selected region of interest, the wireframe model comprising a plurality of vertices, each vertex having a coordinate position;
  - c. refinement means for refining the wireframe model by adjusting the coordinate positions



- of the vertices to more accurately represent the region of interest; and
- d. rendering means for rendering the wireframe model in an interactive three-dimensional display, therein producing a virtual three-dimensional environment.
31. The system of Claim 30 wherein the refinement means adjusts the coordinate positions of vertices of the wireframe model along a normal vector associated with each vertex at a rate that is a function of a surface curvature at the coordinate position.
32. The system of Claim 31 wherein the refinement means adjusts the coordinate positions of the vertices along a normal vector associated with each vertex by determining an inner wall position and an outer wall position an analysis of intensity profiles of voxel values along the normal vector.
33. The system of Claim 30 wherein the modeling means, and the refining means are located remotely from the volume formation means.
34. A system for interactively displaying three dimensional structures comprising:
- a. volume formation means for forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with a three dimensional body;

- b. wireframe producing means for producing a wireframe model of a selected region of interest of the three-dimensional volume;
  - c. deforming means for deforming the wireframe model to more accurately represent the region of interest; and
  - d. rendering means for rendering the wireframe model in an interactive three-dimensional display, therein producing a virtual three-dimensional environment.
35. A system for interactively displaying a three-dimensional rendering of a structure having a lumen and indicating regions having abnormal structure comprising:
- a. volume formation means for forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with the three-dimensional structure;
  - b. wireframe generation means for generating a wireframe model of a selected region of interest;
  - c. vertex grouping means for grouping the vertices of the wireframe model into regions having abnormal structure;
  - d. analyzing means for analyzing the regions having abnormal structure; and
  - e. rendering means for rendering the wireframe model in an interactive three-dimensional display to indicate the regions having abnormal structure.

36. The system of Claim 35 wherein the analyzing means comprises means for evaluating two principal curvatures associated with each point in an elliptical region.
37. The system of Claim 36 wherein the analyzing means comprises a means for identifying ridges.
38. The system of claim 37 comprising wireframe deforming means for deforming the wireframe model to more accurately represent the region of interest.
39. A computer-implemented method for interactively displaying a three-dimensional rendering of a structure having a lumen and indicating regions having abnormal structure comprising the steps of:
- a. forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with the three-dimensional structure;
  - b. generating a wireframe model of a selected region of interest, the wireframe model comprising a plurality of vertices, each vertex having a coordinate position;
  - c. grouping the vertices of the wireframe model into regions having a characteristic indicating abnormal structure; and
  - d. analyzing the regions having abnormal structure to identify regions having a high degree of abnormality.

40. The method of claim 39 comprising the steps of rendering the wireframe model in an interactive three-dimensional display to indicate the regions having abnormal structure.
41. A computer-implemented method for interactively displaying a three-dimensional rendering of a structure having a lumen comprising the steps of:
- a. forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with the three-dimensional structure;
  - b. generating a wireframe model of a selected region of interest;
  - e. deforming the wireframe model to more accurately represent the region of interest; and
  - f. rendering the wireframe model in an interactive three-dimensional display.
42. A computer-implemented method for interactively displaying a three-dimensional rendering of a structure having a lumen comprising the steps of:
- a. forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with the three-dimensional structure;
  - b. generating a wireframe model of a selected area of interest;
  - c. refining the wireframe model of the selected region of interest to more accurately

- represent the region of interest; and
- d. rendering the wireframe model in an interactive three-dimensional display.
43. The method of Claim 42 wherein the refinement step adjusts coordinate positions of vertices of the wireframe model along a normal vector associated with each vertex at a rate that is a function of a surface curvature at the coordinate position.
44. The method of Claim 43 wherein the refinement step adjusts the coordinate positions of the vertices along a normal vector associated with each vertex by determining an inner wall position and an outer wall position from a first and a second degree derivative associated with the coordinate positions of the vertices.
45. A method for guiding biopsy of a tissue by accessing the tissue via a three-dimensional structure having a lumen comprising the steps of:
- a. forming a three-dimensional volume of data from a series of two-dimensional images representing at least one physical property associated with the three-dimensional structure and at least one physical property associated with the tissue;
  - b. generating a wireframe model of a selected region of interest;
  - c. rendering the wireframe model in an interactive three-dimensional display;
  - d. inserting a simulated biopsy needle within

- the lumen of the three-dimensional structure;
- e. determining the position of the simulated biopsy needle within the three-dimensional structure; and
  - f. displaying the rendered, interactive three-dimensional display from a perspective indicating the position of the simulated biopsy needle within the three-dimensional structure.
46. The method of Claim 45 comprising the step of indicating the position of the simulated biopsy needle relative to the position of the tissue.
47. The method of Claim 46 wherein the indicating step comprises the steps of:
- a. generating a zone around the tissue; and
  - b. issuing an indicator when a tip of the biopsy needle is within the zone.
48. The method of Claim 46 wherein the indicating step comprises the step of determining a distance and a trajectory from the simulated biopsy needle to the tissue.
49. The method of Claim 39, 41, 42, or 45 comprising an image acquisition step wherein a series of two-dimensional images representing at least one physical property associated with the structure is acquired, wherein the image acquisition step comprises a patient preparation step.

50. The method of Claim 49 wherein the patient preparation step comprises the step of administering to a patient a nonionic intravenous iodinated contrast bolus.
51. The method of Claim 49 wherein the patient preparation step comprises the step of administering to a patient an oral contrast agent for opacifying stool and water.

0055600 002304

Figure 1

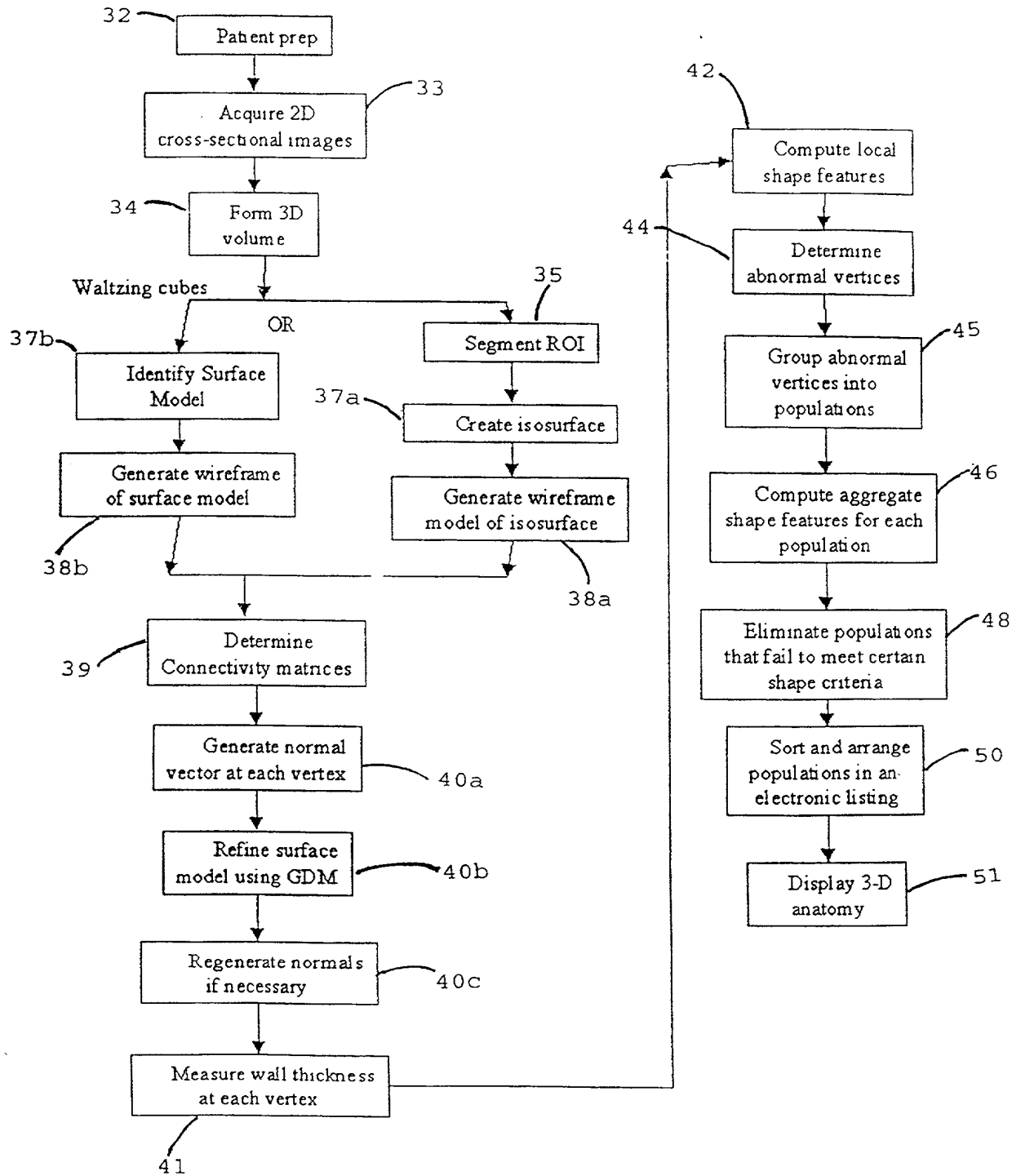
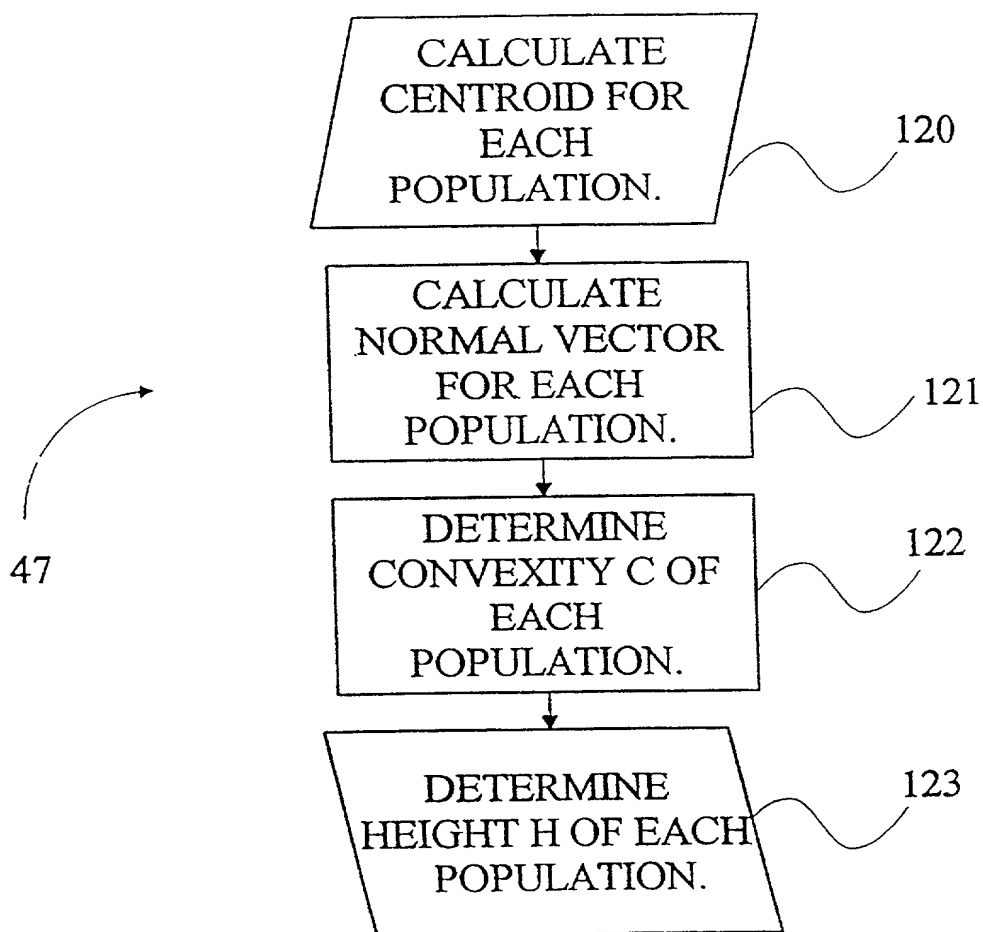




FIG. 2



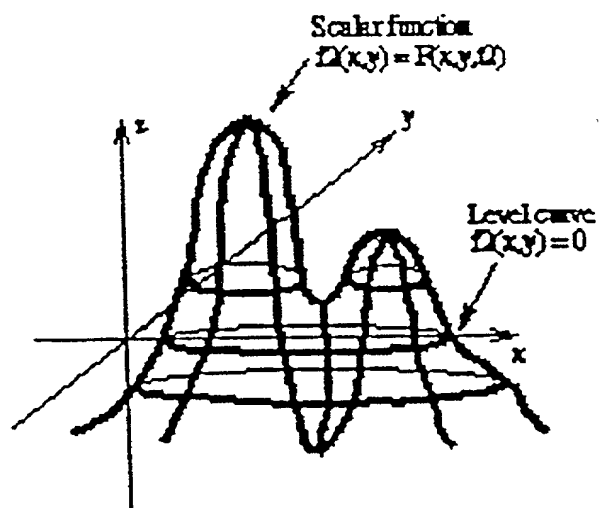
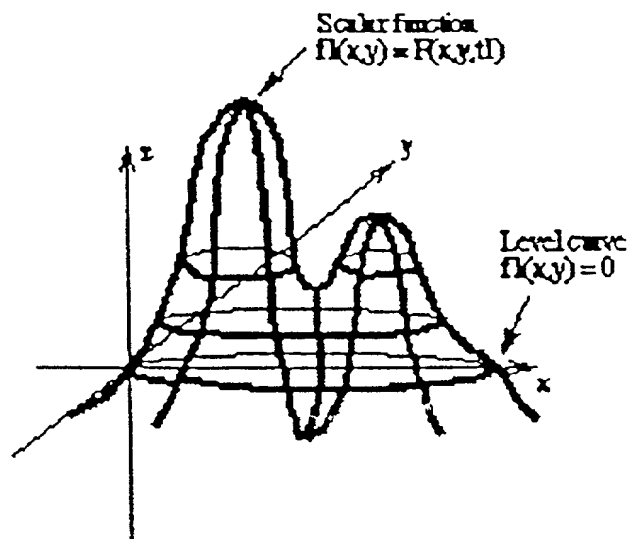


Fig. 3

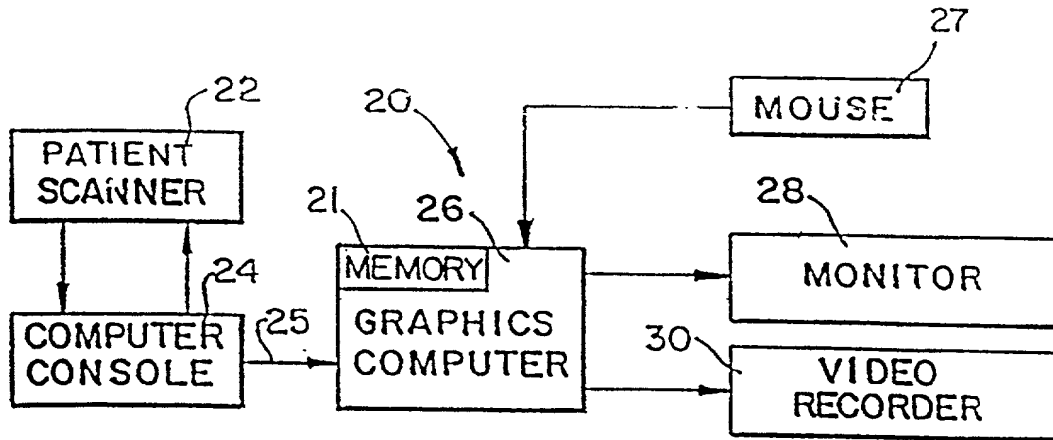
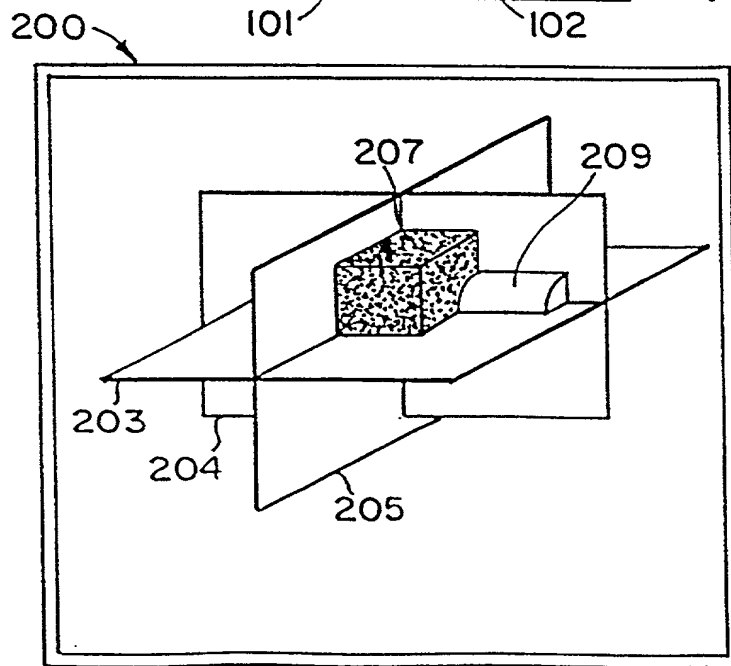
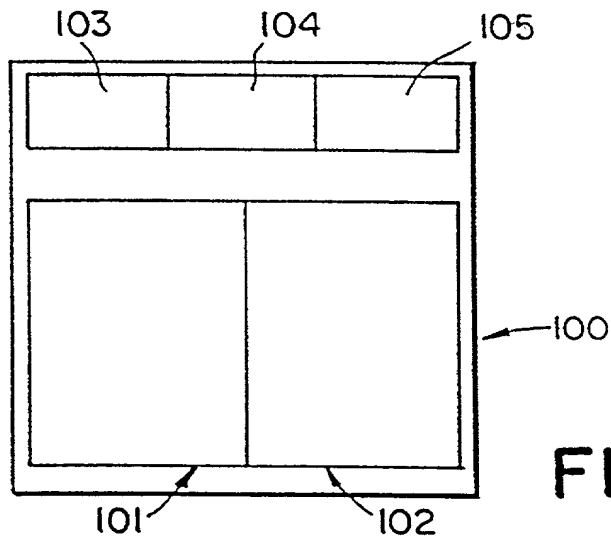
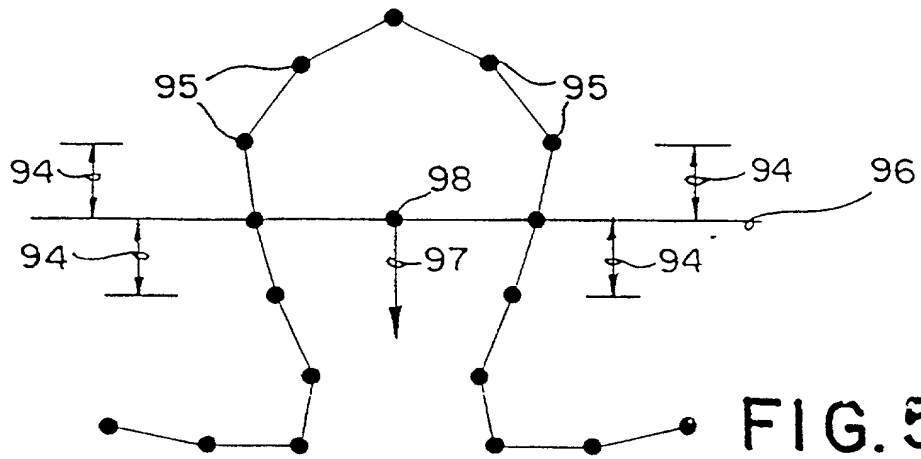
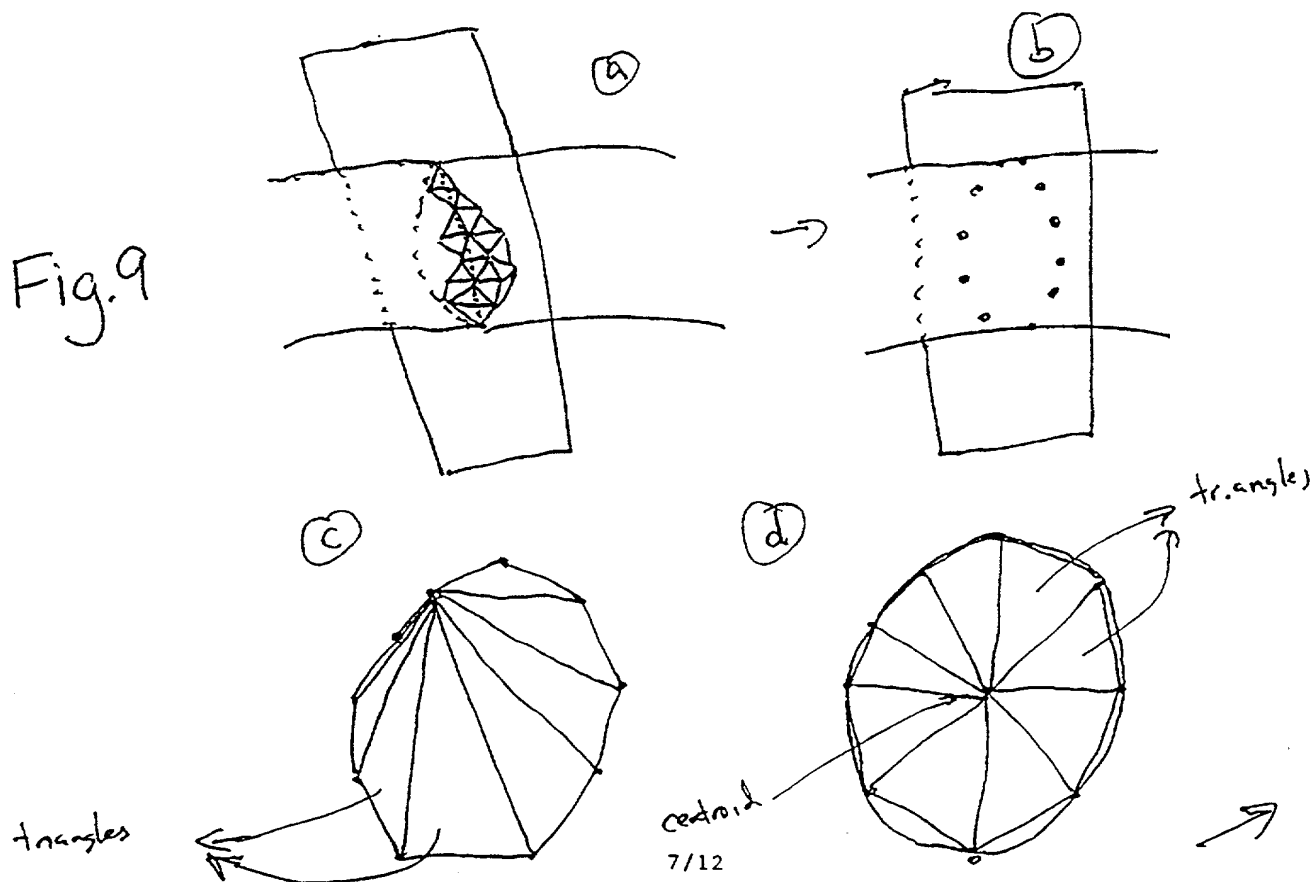
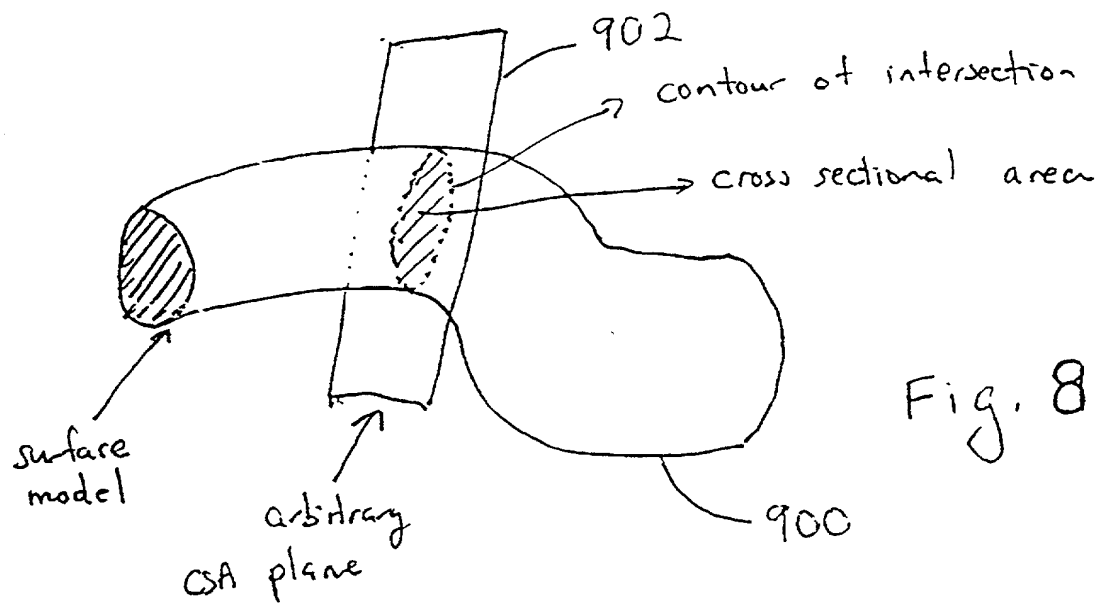


FIG. 4







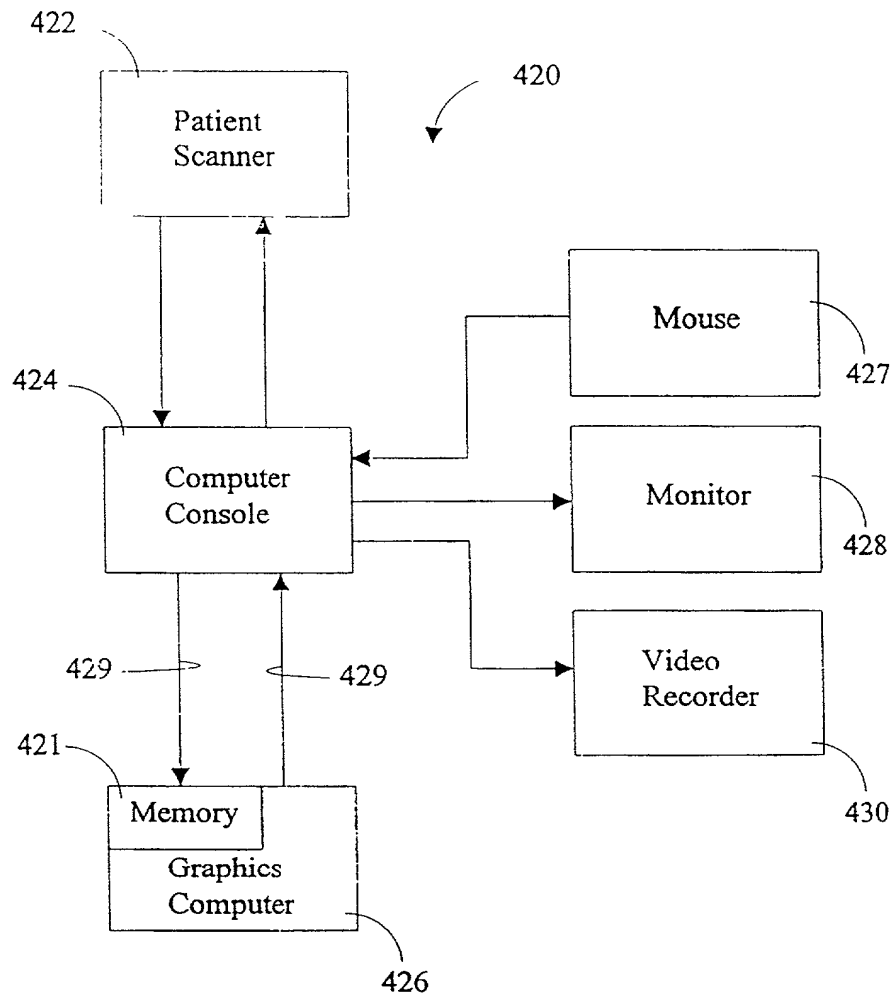


Fig. 11

Figure 12

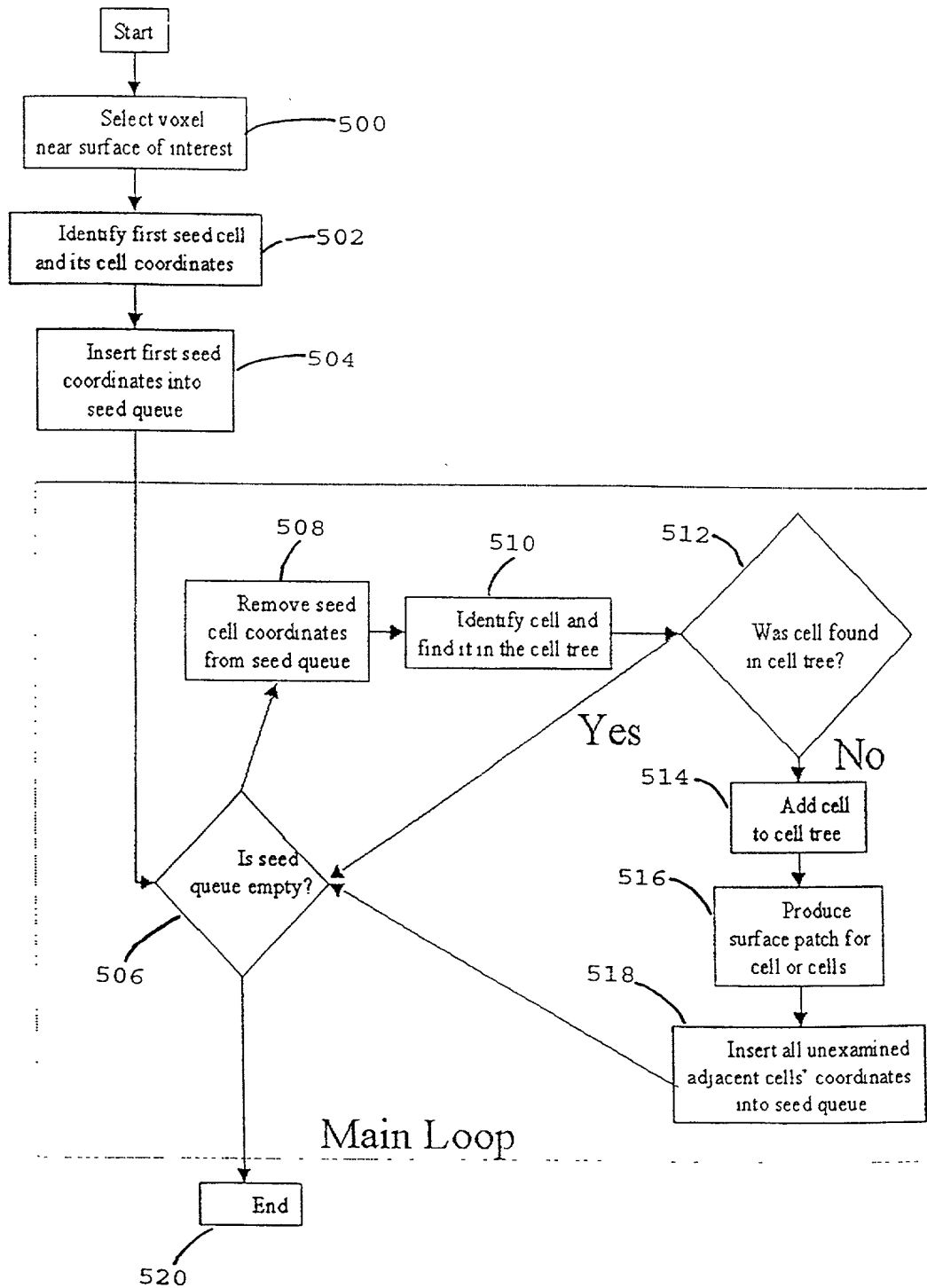




Fig. 13

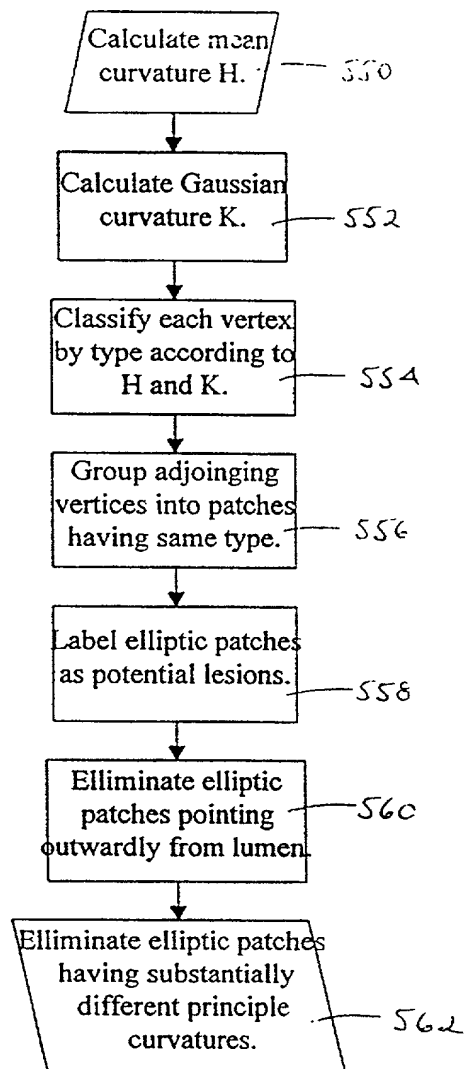
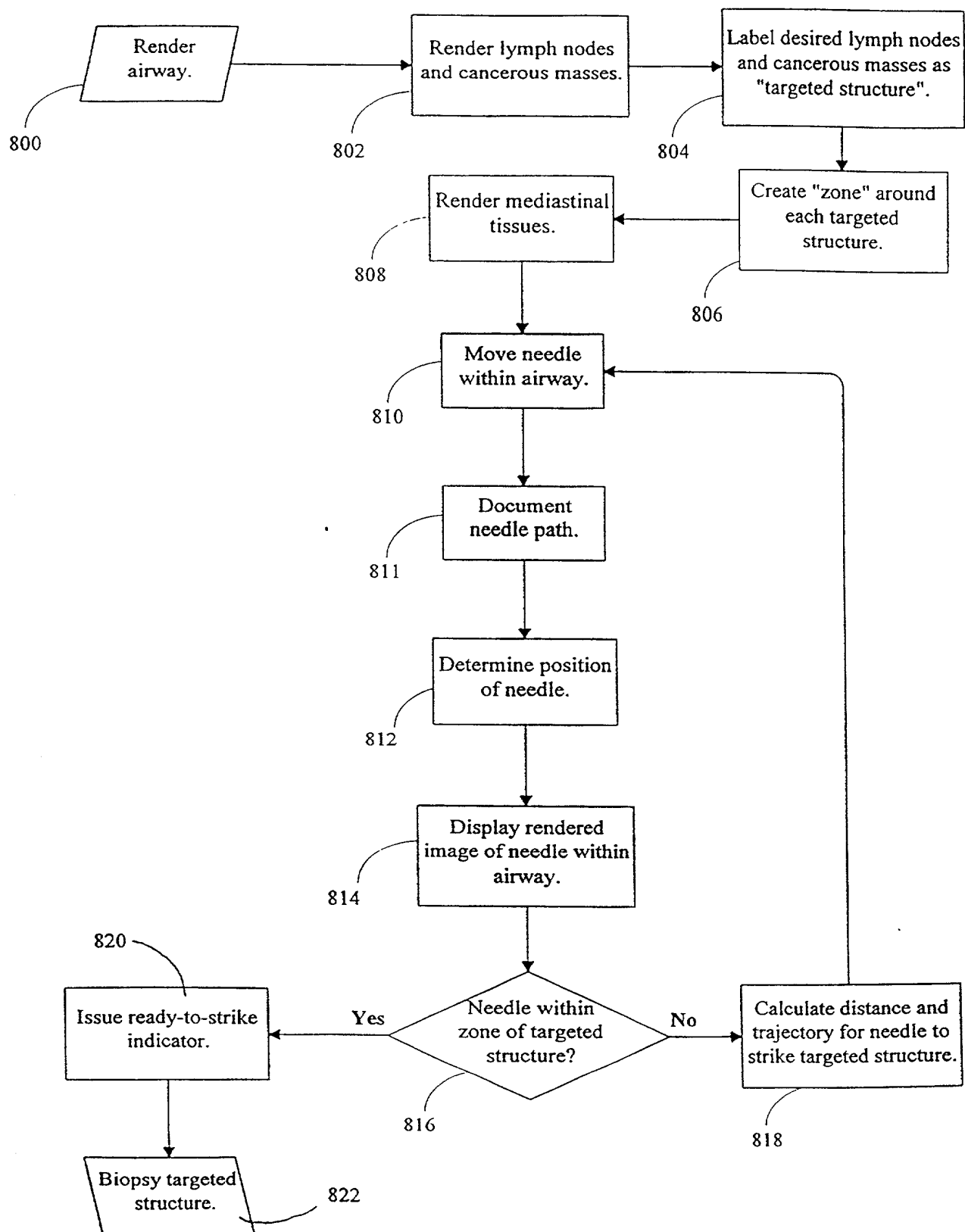


Fig. 14



200	200	0	0	0	0	0	0	0	0
200	200	200	0	0	0	0	0	0	0
200	200	200	200	0	0	0	0	0	0
100	200	200	200	200	0	0	0	0	0
100	100	200	200	200	200	0	0	0	0
100	100	100	200	200	200	200	0	0	0
100	100	100	100	200	200	200	200	0	0
100	100	100	100	100	200	200	200	200	0
100	100	100	100	100	100	200	200	200	200
100	100	100	100	100	100	100	200	200	200

600

Fig. 15

601

200	150	90	0	0	0	0	0	0	0
200	200	150	90	0	0	0	0	0	0
200	200	200	150	90	0	0	0	0	0
120	200	200	200	150	90	0	0	0	0
100	120	200	200	200	150	90	0	0	0
100	100	120	200	200	200	150	90	0	0
100	100	100	120	200	200	200	150	90	0
100	100	100	100	120	200	200	200	150	90
100	100	100	100	100	120	200	200	200	150
100	100	100	100	100	100	120	200	200	200

Fig. 16

0	0	90	0	0	0	0	0	0	0
0	0	0	90	0	0	0	0	0	0
0	0	0	0	90	0	0	0	0	0
120	0	0	0	0	90	0	0	0	0
100	120	0	0	0	0	90	0	0	0
100	100	120	0	0	0	0	90	0	0
100	100	100	120	0	0	0	0	90	0
100	100	100	100	120	0	0	0	0	90
100	100	100	100	100	120	0	0	0	0
100	100	100	100	100	100	120	0	0	0

Fig. 17

#4

## DECLARATION, POWER OF ATTORNEY AND POWER TO INSPECT

As a below named inventor, I hereby declare:

that my residence, post office address and citizenship are as stated below next to my name;

that I verily believe I am the original, first and sole inventor (if only one name is listed below) or an original, first and joint inventor (if plural inventors are named below) of the invention entitled:

the specification of which [check one(s) applicable]

☒ was filed May 23, 2001 as U.S. Application No. 09/856,683  
☒ and was amended by Amendment filed June 25, 2001 (if applicable);  
☒ and was amended by Amendment filed concurrently herewith; [or];  
\_\_\_ is attached to this Declaration, Power of Attorney and Power to Inspect;

that I have reviewed and understand the contents of the above-identified specification, including the claims, as amended by any amendment referred to above; and

that I acknowledge my duty to disclose information which is material to the examination of this application in accordance with Rule 56(a) [37CFR§1.56(a)].

**CLAIM UNDER 35 USC §119(e):** I hereby claim the benefit under 35 USC §119(e) of the prior United States application(s) listed below:

**Provisional Application No.**

60/109,966

**Filing Date**

**Day/Mo/Year**

November 25, 1998

**CLAIM UNDER 35 USC §120:** I hereby claim the benefit under 35 USC §120 of the prior United States application(s) listed below:

**Prior U.S. Application(s)**

09/299,061

08/805,584

**Filing Date**

**Day/Mo/Year**

April 23, 1999

February 24, 1997

**Status**

**Pending-Patented-Abandoned**

Pending

Patent No. 5,920,319

Insofar as the subject matter of each of the claims of this application is not disclosed in the prior United States application in the manner provided by the first paragraph of 35 USC §112, I acknowledge the duty to disclose material information as defined in Rule 56(a) [37 CFR §1.56(a)] which occurred between the filing date of the prior U.S. application and the national or PCT international filing date of this application.

**POWER OF ATTORNEY:** As inventor, I hereby appoint **DANN, DORFMAN, HERRELL AND SKILLMAN, P.C.** of Philadelphia, PA, and the following individual(s) as my attorneys or agents with full power of substitution to prosecute this application and to transact all business in the Patent and Trademark Office connected therewith: **Donald R. Piper, Jr. Reg. No. 29,337** and **Vincent T. Pace, Reg. No. 31,049.**

**POWER TO INSPECT:** I hereby give **DANN, DORFMAN, HERRELL AND SKILLMAN, P.C.** of Philadelphia, PA or its duly accredited representatives power to inspect and obtain copies of the papers on file relating to this application.

**SEND CORRESPONDENCE TO: CUSTOMER NO. 000110**

**DIRECT INQUIRIES TO: DONALD R. PIPER, JR.; Telephone (215) 563-4100/Facsimile (215) 563-4044**

I hereby declare that all statements made herein of my own knowledge are true and that all statements made on information and belief are believed to be true; and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful false statements may jeopardize the validity of the application or any patent issued thereon.

**SOLE OR FIRST JOINT INVENTOR**

Full Name DAVID J. VINING  
First Middle Last

Signature [Signature]

Date 8/15/01

Residence Winston-Salem North Carolina

City State or Country NC

Citizenship United States of America

Post Office Address:

1725 Briar Lake Road

Winston-Salem, North Carolina 27103

City State or Country Zip Code

**SECOND JOINT INVENTOR (if any)**

Full Name GORDON W. HUNT  
First Middle Last

Signature [Signature]

Date 8/16/01

Residence Winston-Salem North Carolina

City State or Country NC

Citizenship United States of America

Post Office Address:

608 Shadylawn Court

Winston-Salem, North Carolina 27104

City State or Country Zip Code

3-00

**THIRD JOINT INVENTOR (if any)**

Full Name DAVID K. AHN  
First Middle Last  
Signature [Signature]  
Date 8/14/01  
Residence Greensboro, North Carolina  
City State or Country NC  
Citizenship United States of America  
Post Office Address:  
2005 Tennyson Drive 5026 Mereworth Ct  
Greensboro, North Carolina 27409  
City State or Country Zip Code  
Winston-Salem, NC 27104

5-00

**FIFTH JOINT INVENTOR (if any)**

Full Name YAORONG GE  
First Middle Last  
Signature [Signature]  
Date 8/14/01  
Residence Winston-Salem, North Carolina  
City State or Country NC  
Citizenship China U.S.A.  
Post Office Address:  
305 Rockmont Drive  
Winston-Salem, North Carolina 27104  
City State or Country Zip Code

**FOURTH JOINT INVENTOR (if any)**

Full Name DAVID R. STELTS  
First Middle Last  
Signature [Signature]  
Date 8-16-2001  
Residence Silver Creek Georgia Rock Hill, SC  
City State or Country SC  
Citizenship United States of America  
Post Office Address:  
762 Shifflett Road  
Silver Creek, Georgia 30173  
City State or Country Zip Code  
1966 Steeplechase Dr  
Rock Hill, SC  
29732